

95698/

## SEARCH REQUEST FORM

RECEIVED

96062

Scientific and Technical Information Center

JUN -3 2003

Requester's Full Name Jeffrey E. Russel Examiner # 62785 Date 6-3-2003  
 An Unit 1654 Phone Number 308-3975 Serial Number 5097815978  
 Mail Box and Bldg Room Location \_\_\_\_\_ Results Format Preferred (circle) PAPER DISK E-MAIL  
CM1-11D13/CM1-9807

If more than one search is submitted, please prioritize searches in order of need.

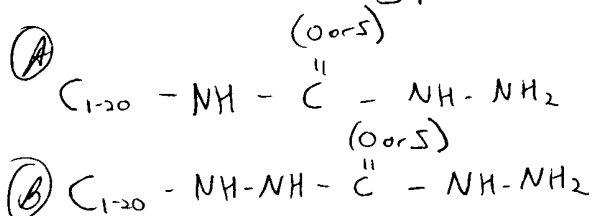
\*\*\*\*\*  
 Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc., if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention Hydrazine-Based And Carbonyl-Based Bifunctional Crosslinking Reagents  
 Inventors (please provide full names) D. Schwartz

Earliest Priority Filing Date 3-22-2001

\*For Sequence Searches Only\* Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

Please search the following partial structures:



Keywords are crosslink?, bifunctional, heterobifunctional, immobility?, conjugate?.

Thank you.

JER

## STAFF USE ONLY

## Type of Search

## Vendors and cost where applicable

Searcher _____	NA Sequence (#) _____	STN _____
Searcher Phone # _____	AA Sequence (#) _____	Dialog _____
Searcher Location _____	Structure (#) <u>2</u>	Quick Ortho _____
Date Received <u>6/4/03</u>	Bibliographic _____	Indexing _____
Date Indexed <u>6/6/03</u>	Integration _____	Index Notes _____
Searcher Prep & Review Time <u>20</u> / <u>10</u>	Fulltext _____	Sequence Systems _____
Client Prep Time _____	Patent Family _____	ANALYSIS/INTERPRET _____
_____ <u>30</u> / <u>10</u>	Other _____	Other Services _____

=&gt; d ilib aks hitstr 14 1-5

L4 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2003 APS  
 ACCESSION NUMBER: 2001:55626 HCAPLUS  
 DOCUMENT NUMBER: 137:115644  
 TITLE: Ternary biomolecule/polymer/surface-based  
 immobilization methods  
 INVENTOR(S): Schwartz, David A.  
 PATENT ASSIGNEE(C): USA  
 SOURCE: PCT Int. Appl., 13 pg.  
 CODEN: PINKDC  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001-057412	A2	20020715	WO 2001-051161	20020116
WO 2001-057412	A3	20020217		

WI: AL, AM, AT, AU, AZ, BA, BE, BG, BF, BY, CA, CH, CN, CU, CZ, DE,  
 DK, EE, ES, FI, GB, GR, GE, GH, GM, HF, HU, IL, IN, IS, JP,  
 KE, KG, KP, KR, KZ, LC, LK, LF, LS, LT, LU, LV, ML, MG, MK, MN,  
 MW, MX, ND, NE, NL, NO, NZ, OM, OS, PA, PE, PG, PH, PK, PL, PT, RU, TJ, TM,  
 TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AE, BY, KG, KZ, MD, RU, TJ, TM  
 FW: GH, GM, KE, LS, MW, MC, SD, SL, SC, TG, UG, ZM, ZW, AT, BE, CH,  
 CY, DE, DK, ES, FI, FR, GE, GR, IE, IT, LU, MC, NL, PT, SE, TR,  
 BF, BJ, CF, CG, CI, CM, GA, GU, GQ, GW, ML, MF, NE, NG, SD, TD, TG

US 2001146504 A1 20021910 US 2001-50277 20030115  
 PRIORITY APPLN. INFO.: US 2001-262094P P 20010116  
 US 2002-50277 A 20020115  
 US 2000-191136P P 20000322

AB: Immobilizing natural or synthetic biomols. onto surfaces comprises  
 covalently linking the natural or synthetic biomol. to a mono- or  
 bi-functional polymer and covalently and/or electrostatically immobilizing  
 the biomol./polymer conjugate to an unmodified or modified surface, where  
 the biomol. is an oligonucleotide, a polynucleotide, a protein, a  
 glycoprotein, a peptide or a carbohydrate that was modified to incorporate  
 .gtoreq.1 nucleophilic groups comprising an aliph. or arom. amino, thiol,  
**hydrazine**, thiosemicarbazide, hydrazide, thiocarbamide, carbamide,  
 aminoxyl, a deriv. of 2-hydrazinopyridine or aminoxylacetic acid or  
 .gtoreq.1 electrophilic groups comprising an aliph. or arom. aldehyde,  
 ketone, epoxide, isocyanate, isothiocyanate, succinimidyl ester or  
 cyanamic chloride or a linkable arom. aldehyde or ketone and the surface  
 was modified to possess either neutral, cationic or anionic groups or a  
 combination neutral, anionic and/or cationic moieties.

IT: **25104-18-1DP**, Polylysine, reaction products with succinimidyl  
 hydrazinonicotinate acetone hydrate, conjugates **38000-06-5DP**,  
 Polylysine, reaction products with succinimidyl hydrazinonicotinate  
 acetone hydrate, conjugates **60444-78-2DP**, Succinimidyl  
 4-formylbenzoate, reaction products with polylysine, conjugates with  
 oligonucleotides **362522-50-7DP**, Succinimidyl  
 (5-hydrazinonicotinate acetone hydrate, polymer deriv., conjugates with  
 oligonucleotides  
 EL: BIOL (Biological use, unclassified); IMF (Industrial manufacture); BIOL  
 (Biological study); PREP (Preparation); USES (Uses)  
 (ternary biomol./polymer/surface-based immobilization systems)

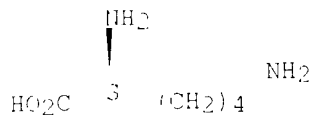
RI: 25104-18-1 HCAPLUS  
 CN: L-Lysine, homopolymer (PT) (CA INDEX NAME)

CM 1

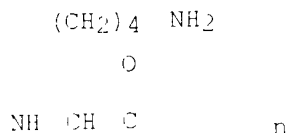
CPN 56-87-1

CMF C6 H14 N2 O2

Absolute stereochemistry.

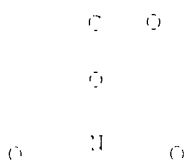


RN 38006-06-5 HCAPLUS  
CN Poly[imino[(1S)-1-(4-aminobutyl)-2-oxo-1,2-ethanediyl]] (9CI) (CA INDEX NAME)

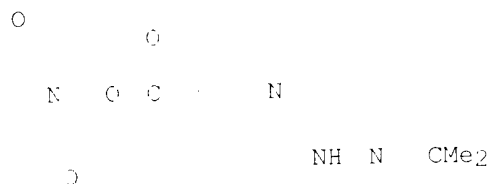


RN 60444-78-2 HCAPLUS  
CN Benzaldehyde, 4-[[[(2,5-dioxo-1-pyrrolidinyl)oxy]carbonyl]- (9CI) (CA INDEX NAME)

CHO



RN 62522-50-7 HCAPLUS  
CN 1,5-Pyrrolidinedione, 1-[[[1,6-dihydro-6-[(1-methylethylidene)hydrazono]-3-pyridinyl]carbonyl]oxy]- (9CI) (CA INDEX NAME)



L4 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMEER: 2001:713345 HCAPLUS

DOCUMENT NUMBER: 135:172864

TITLE: **Hydrazine**-based and carbonyl-based bifunctional crosslinking reagents for biomolecules, drugs, and synthetic polymersINVENTOR(S): **Schwartz, David A.**

PATENT ASSIGNER(S): Celulink, Inc., USA

SOURCE: PCT Int. Appl., 47 pp.

CODEN: PEXEDL

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001070685	A2	20010917	WO 2001-039252	20010122
WO 2001070685	A3	20030327		
W:	AE, AG, AL, AM, AT, AU, AV, BA, BB, BC, BD, BE, BF, BG, BH, BI, BJ, BK, BL, BM, BN, BO, BP, BQ, BR, BS, BT, BU, BV, BW, BX, BY, BZ, CA, CB, CC, CD, CE, CF, CG, CH, CI, CJ, CK, CL, CM, CN, CO, CP, CQ, CR, CS, CT, CU, CV, CW, CX, CY, CZ, DA, DB, DC, DD, DE, DF, DG, DH, DI, DJ, DK, DL, DM, DN, DO, DP, DQ, DR, DS, DT, DU, DV, DW, DX, DY, DZ, EA, EB, EC, ED, EE, EF, EG, EH, EI, EJ, EK, EL, EM, EN, EO, EP, EQ, ER, ES, ET, EU, EV, EW, EX, EY, EZ, FA, FB, FC, FD, FE, FF, FG, FH, FI, FJ, FK, FL, FM, FN, FO, FP, FQ, FR, FS, FT, FU, FV, FW, FX, FY, FZ, GA, GB, GC, GD, GE, GF, GH, GI, GJ, GK, GL, GM, GN, GO, GP, GQ, GR, GS, GT, GU, GV, GW, GX, GY, GZ, HA, HB, HC, HD, HE, HF, HG, HH, HI, HJ, HK, HL, HM, HN, HO, HP, HQ, HR, HS, HT, HU, HV, HW, HX, HY, HZ, IA, IB, IC, ID, IE, IF, IG, IH, II, IJ, IK, IL, IM, IN, IO, IP, IQ, IR, IS, IT, IU, IV, IW, IX, IY, IZ, JA, JB, JC, JD, JE, JF, JG, JH, JI, JJ, JK, JL, JM, JN, JO, JP, JQ, JR, JS, JT, JU, JV, JW, JX, JY, JZ, KA, KB, KC, KD, KE, KF, KG, KH, KI, KJ, KK, KL, KM, KN, KO, KP, KQ, KR, KS, KT, KU, KV, KW, KX, KY, KZ, LA, LB, LC, LD, LE, LF, LG, LH, LI, LJ, LK, LM, LN, LO, LP, LQ, LR, LS, LT, LU, LV, LW, LX, LY, LZ, MA, MB, MC, MD, ME, MF, MG, MH, MI, MJ, MK, ML, MN, MO, MP, MQ, MR, MS, MT, MU, MV, MW, MX, MY, MZ, NA, NB, NC, ND, NE, NF, NG, NH, NI, NJ, NK, NL, NM, NO, NP, NQ, NR, NS, NT, NU, NV, NW, NX, NY, NZ, OA, OB, OC, OD, OE, OF, OG, OH, OI, OJ, OK, OL, OM, ON, OO, OP, OQ, OR, OS, OT, OU, OV, OW, OX, OY, OZ, PA, PB, PC, PD, PE, PF, PG, PH, PI, PJ, PK, PL, PM, PN, PO, PP, PQ, PR, PS, PT, PU, PV, PW, PX, PY, PZ, QA, QB, QC, QD, QE, QF, QG, QH, QI, QJ, QK, QL, QM, QN, QO, QP, QQ, QR, QS, QT, QU, QV, QW, QX, QY, QZ, RA, RB, RC, RD, RE, RF, RG, RH, RI, RJ, RK, RL, RM, RN, RO, RP, RQ, RR, RS, RT, RU, RV, RW, RX, RY, RZ, SA, SB, SC, SD, SE, SF, SG, SH, SI, SJ, SK, SL, SM, SN, SO, SP, SQ, SR, SS, ST, SU, SV, SW, SX, SY, SZ, TA, TB, TC, TD, TE, TF, TG, TH, TI, TJ, TK, TL, TM, TN, TO, TP, TQ, TR, TS, TU, TV, TW, TX, TY, TZ, UA, UB, UC, UD, UE, UF, UG, UH, UI, UJ, UK, UL, UM, UN, UO, UP, UQ, UR, US, UT, UV, UW, UX, UY, UZ, VA, VB, VC, VD, VE, VF, VG, VH, VI, VJ, VK, VL, VM, VN, VO, VP, VQ, VR, VS, VT, VU, VV, VW, VX, VY, VZ, WA, WB, WC, WD, WE, WF, WG, WH, WI, WJ, WK, WL, WM, WN, WO, WP, WQ, WR, WS, WT, WU, WV, WW, WX, WY, WZ, XA, XB, XC, XD, XE, XF, XG, XH, XI, XJ, XK, XL, XM, XN, XO, XP, XQ, XR, XS, XT, XU, XV, XW, XX, XY, XZ, YA, YB, YC, YD, YE, YF, YG, YH, YI, YJ, YK, YL, YM, YN, YO, YP, YQ, YR, YS, YT, YU, YV, YW, YX, YZ, ZA, ZB, ZC, ZD, ZE, ZF, ZG, ZH, ZI, ZJ, ZK, ZL, ZM, ZN, ZO, ZP, ZQ, ZR, ZS, ZT, ZU, ZV, ZW, ZX, ZY, ZZ			
US 2003013857	A1	20030116	US 2001 813973	20010321
EP 1215699	A2	20030604	EP 2001-029666	20010321
F:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, PO, MK, CY, AL, TR			
PRIORITY APPLN. INFO.:			US 2000-191868	F 20000321
			WO 2001-039252	W 20010321

OTHER SOURCE(S): MAPPAT 135:172864

AB F-agents and methods are provided for bifunctional crosslinking and immobilizing biomols., drugs, and synthetic polymers. The reagents of formula  $\text{BRANHNH}_2\text{bul.HX}$  [wherein A =  $\text{NHCO}$ ,  $\text{NHCS}$ ,  $\text{NHNHCO}$ ,  $\text{NHNHCS}$ , or a direct bond; B = an amino or thio reactive moiety; F = specified aliph. divalent groups contg. any combination of cycloalkylene, C(R10)2, CR10:CR10, C:CR12R13, CR12F13, C:triplond.C, C, SGa, NR10, N-R12R13, CL, etc.; a = 0-1; b = 0-3; G = O or NR10; L = S, O, or NR10; R10 = specified monovalent groups; R12 and R13 = independently H, (cyclo)alkyl, alkenyl, alkynyl, or (hetero)aryl; or R12 and R13 together form (cyclo)alkylene or alkenylene; X = neg. counterion; or a deriv. thereof] possess a thiol or amino reactive group and a hydrazine or oxoamine moiety. Conjugates and immobilized biomols. are also provided. For example, hydratinicotinic acid was converted to the acetone hydrazine and treated with N-hydroxysuccinimide to give the crosslinking agent, succinimidyl O-hydratinicotinate acetone hydrazine (I), in 33% yield. A soln. of ovalbumin in PBS and EDTA was added to a soln. of I in DMF and the mixt. incubated at room temp. for 4 h to afford the **hydrazine**-modified protein, which exhibited a molar extinction coeff. of 22,000 at 360 nm.

IT 60444-78-2 362522-64-3

FL: RCT (Reactant); FACT (Reactant + reagent)

crosslinking agent; prepn. of **hydrazine**- and carbonyl-based bifunctional crosslinking agents and use with biomols., drugs, and synthetic polymers)

RN 60444-78-2 HCAPLUS

CN Benzaldehyde, 4-[[[(2,5-dioxo-1-pyrrolidinyl)oxy]carbonyl]- (9CI) (CA INDEX NAME)

CHO

C O

O

O N O

RN 362522-64-3 HCAPLUS

CN Hydrazinecarboxamide, N-[4-[[[(2,5-dioxo-1-pyrrolidinyl)oxy]carbonyl]phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)

O

H<sub>2</sub>N NH C NH

C O

O

O N O

● HCl

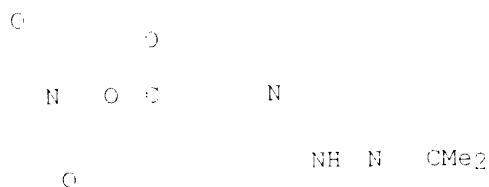
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362522-53-0P 362522-54-1P 362522-55-2P  
362522-56-3P 362522-57-4P 362522-58-5P

EL: PCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); FACT (Reactant or reagent)

(crosslinking agent; prepn. of **hydrazine-** and carbonyl-based bifunctional crosslinking agents and use with biomols., drugs, and synthetic polymers)

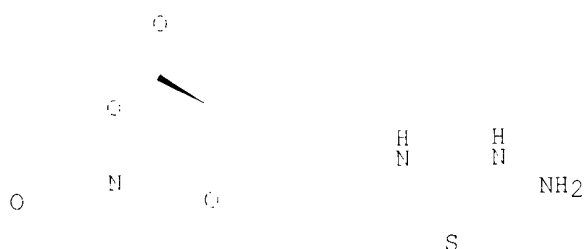
RN 362522-50-7 HCAPLUS

CN 2,5-Pyrrolidinedione, 1-[[[1,6-dihydro-6-[(1-methylethylidene)hydrazono]-3-pyridinyl]carbonyl]oxy]- (9CI) (CA INDEX NAME)



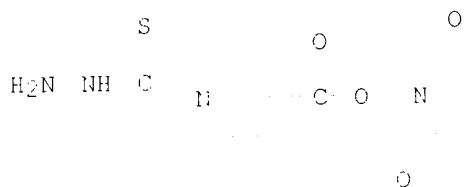
RN 362521-51-8 HCAPLUS  
 CN Hydrazinecarbothioamide, N-[[trans-4-[[[(2,5-dioxo-1-pyrrolidinyl)oxy]carbonyl]cyclohexyl]methyl]-, monohydrochloride (9CI)  
 (CA INDEX NAME)

Relative stereochemistry.



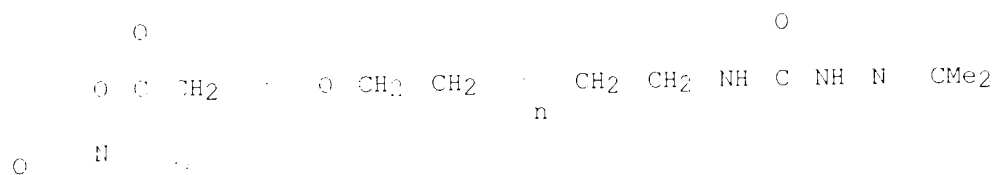
● HCl

RN 362522-52-9 HCAPLUS  
 CN 1-Pyrrolidinecarboethioic acid, 3-[[[(2,5-dioxo-1-pyrrolidinyl)oxy]carbonyl]-, hydrazide, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

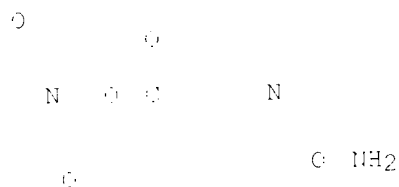
RN 362522-93-0 HCAPLUS  
 CN Ecoly(oxy-1,2-ethanediyl), .alpha.-[2-[(2,5-dioxo-1-pyrrolidinyl)oxy]-2-oxoethyl]-.omega.-[2-[[[(1-methylethylidene)hydrazino]carbonyl]amino]ethyl]- (9CI) (CA INDEX NAME)



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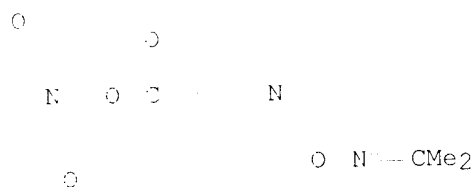
FN      16252.-54-1  HCAPLUS
CN      2,5-Pyrrolidinedione, 1-[[[6-(aminoxy)-3-pyridinyl]carbonyl]oxy]-,
monohydrochloride (HCl)  (CA INDEX NAME)

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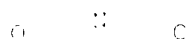
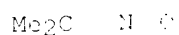


● HCl

RN 162520-55-2 HCAFLUS  
 CN 2,5-Pyrrolidinedione, 1-[[[6-[[[(1-methylethylidene)amino]oxy]-3-pyridinyl]carbonyl]oxy]- (9CI) (CA INDEX NAME)



RN 36250-55-3 HCAPLUS  
CN 1H-Pyrrole-2,5-dione, 1-[4-[[ (1-methylethylidene)amino]oxy]phenyl]- (9CI)  
(CA INDEX NAME)



FN 3625-2-57-4 HCAPLUS

OE: O

$$\text{EtO} \quad \text{Si} \quad (\text{CH}_2)_3 \quad \text{NH} \quad \text{C}$$

○E=

$$\text{NH} \quad \text{N} \quad \text{CMe}_2$$

FN 362522-58-5 HCAPLUS

FN 362522-58-5 HCAPLUS  
CN 3-Pyridinecarboxamide, N,N'-(aithiodi-2,1-ethanediyl)bis[6-hydrazino-,  
dihydrochloride (9CI) (CA INDEX NAME)

PAGE 1-A

$$\text{N} \quad \text{O} \quad \text{O} \quad \text{N}$$

$$\text{C} \quad \text{NH} \quad \text{CH}_2 \quad \text{CH}_2 \quad \text{S} \quad \text{S} \quad \text{CH}_2 \quad \text{CH}_2 \quad \text{NH} \quad \text{C}$$

NH

$$\text{H}_2\text{N} \quad \text{NE}$$
 $\bullet 2 \text{ HCl}$ 

PAGE 1-B

 $\text{NH}_2$ 

IT 302-01-2DP, **Hydrazine**, derivs., preparation  
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
 (Reactant or reagent)  
 (crosslinking agents; prepn. of **hydrazine**- and carbonyl-based  
 bifunctional crosslinking agents and use with biomols., drugs, and  
 synthetic polymers)

RN 302-01-2 HCAFLUS

RN 302-91-2 HCAFNOS  
CN Hydrazine (7CI, 8CI, 9CI) (CA INDEX NAME)

$$\text{H}_2\text{N} \quad \text{NH}_2$$

IT 6066-82-6, N-Hydroxysuccinimide 25104-18-1,  
Poly-L-lysine 38000-06-5, Poly-L-lysine 133081-24-0,  
6-Hydrazinonicotinic acid 363633-70-9  
RL: RCT (Reactant); RACT (Reactant or reagent)  
(prepn. of **hydrazine**- and carbonyl-based bifunctional  
crosslinking agents and use with biomols., drugs, and synthetic



polymers)  
 EN 6066-82-5 HCAPLUS  
 CN 2,5-Pyrrolidinedione, 1-hydroxy- (9CI) (CA INDEX NAME)

OH

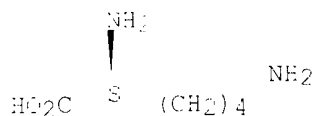
O N O

EN 25104-18-1 HCAPLUS  
 CN L-Lysine, homopolymer (9CI) (CA INDEX NAME)

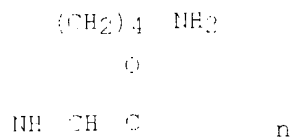
CM 1

CRN 56-87-1  
 CMF C6 H14 N2 O2

Absolute stereochemistry.



EN 38000-06-5 HCAPLUS  
 CN Poly[[imino[(1S)-1-(4-aminobutyl)-2-oxo-1,2-ethanediyl]] (9CI) (CA INDEX NAME)



EN 133081-24-0 HCAPLUS  
 CN 3-Pyridinecarboxylic acid, 6-hydrazino- (9CI) (CA INDEX NAME)

H<sub>2</sub>N NH N

CO<sub>2</sub>H

EN 363633-70-9 HCAPLUS  
 CN DNA, d(T-T-T-T-T-T-T-A-G-C-C-T-A-A-C-T-G-A-T-G-C-C-A-T-G), 5'-(6-aminohexyl hydrogen phosphate) (9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

IT 25104-18-1DP, Poly-L-lysine, hydrazinonicotinamide modified  
 38000-06-5DP, Poly-L-lysine, hydrazinonicotinamide modified  
 364163-70-2P  
 FL: SEN (Synthetic preparation); PREP (Preparation)

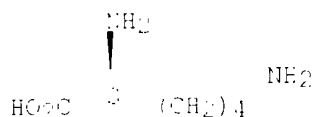
(prepn. of **hydrazine-** and carbonyl-based bifunctional crosslinking agents and use with biomols., drugs, and synthetic polymers)

RN 25104-18-1 HCAPLUS  
CN L-Lysine, homopolymer (9CI) (CA INDEX NAME)

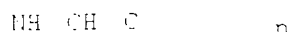
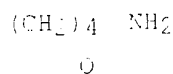
CM 1

CFN 56-37-1  
CMF C6 H14 N2 O2

Absolute stereochemistry.



RN 36060-96-5 HCAPLUS  
CN Poly[amino[(1S)-1-(4-aminobutyl)-2-oxo-1,2-ethanediyl]] (9CI) (CA INDEX NAME)



RN 364163-70-2 HCAPLUS  
CN DNA, 3(T-T-T-T-T-T-A-G-C-C-T-A-A-C-T-G-A-T-G-C-C-A-T-G), 5'-(6-[4-formylbenzoyl]amino)hexyl hydrogen phosphate] (9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

L4 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1991:602039 HCAPLUS

DOCUMENT NUMBER: 115:202039

TITLE: Preparation of hydrazine-modified proteins and their use for the synthesis of technetium-99m-protein conjugates

AUTHOR(S): **Schwartz, David A.**; Abrams, Michael J.; Hauser, Marguerite M.; Gaul, Forrest E.; Larsen, Scott K.; Fauh, Donald; Zubieta, Jon A.

CORPORATE SOURCE: Johnson Matthey Pharm. Res., West Chester, PA, 19380-1407, USA

SOURCE: Bioconjugate Chemistry (1991), 2(5), 333-6  
CODEN: BOCHEH; ISSN: 1043-1802

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The syntheses and protein linking properties of succinimidyl 4-hydrazinobenzoate hydrochloride (SHBH) and succinimidyl 6-hydrazinonicotinate hydrochloride (SHNH), two new heterobifunctional linkers which lead to hydrazine-modified proteins, are described. SHBH-modified proteins are unstable due to the presence of the **phenylhydrazine** moiety. This problem was overcome by synthesizing the hydrazinopyridine analog SHNH, and the conjugates derived from this

linker are stable. Tc(V) cxo precursors readily add to hydrazinopyridine-modified proteins to yield the desired <sup>99m</sup>Tc-radiolabeled protein. <sup>99m</sup>Tc-hydrazinopyridine-polyclonal IgG conjugates are useful agents for the imaging of focal sites of infection.

IT 6066-82-6, N-Hydroxysuccinimide  
 FL: RCT (Reactant); FACT (Reactant or reagent)  
 (esterification of, with hydrazinobenzoic acid deriv.)  
 RN 6066-82-6 HCAPLUS  
 CN 2,5-Pyrrolidinedione, 1-hydroxy- (9CI) (CA INDEX NAME)

OH

O H O

L4 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1991:181413 HCAPLUS

DOCUMENT NUMBER: 114:181413

TITLE: Technetium-99m-human polyclonal IgG radiolabeled via the hydrazine nicotinamide derivative for imaging focal sites of infection in rats

AUTHOR(S): Abrams, Michael J.; Juweid, Malik; TenKate, Caroline I.; Schwartz, David A.; Hausel, Marguerite M.; Gaul, Forrest E.; Fucello, Anthony J.; Rubin, Robert H.; Strauss, H. William; Fischman, Alan J.

CORPORATE SOURCE: Dep. Radiol., Massachusetts Gen. Hosp., Boston, MA, USA

SOURCE: Journal of Nuclear Medicine (1990), 31(12), 2022-8  
 CODEN: JNMEDQ; ISSN: 0161-5595

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The biol. behavior of human polyclonal IgG radiolabeled with <sup>99m</sup>Tc, by a novel method, via a nicotinyl **hydrazine** deriv., was evaluated in rats. Technetium-99m- and indium-111-IgG were coadministered to normal rats and biodistribution was detd. at 2, 6, and 16 h. The inflammation imaging properties of the 2 reagents were compared in rats with deep-thigh infection due to Escherichia coli. Blood clearance of both antibody preps. was well described by a biexponential function: (<sup>99m</sup>Tc-IgG: t1/2 = 3.32 and 57.52 h, <sup>111</sup>In-IgG: 3.93 and 40.71 h). Biodistributions in the solid organs were similar; however, small but statistically significant differences were detected: <sup>99m</sup>Tc-IgG > <sup>111</sup>In-IgG in lung, liver, and spleen; <sup>99m</sup>Tc-IgG < <sup>111</sup>In-IgG in kidney and skeletal muscle. At all 3 imaging times, target-to-background ratio and percent residual activity for the 2 compds. were remarkably similar. These studies establish that human polyclonal IgG labeled with <sup>99m</sup>Tc via a nicotinyl **hydrazine** modified intermediate is equiv. to <sup>111</sup>In-IgG for imaging focal sites of infection in exptl. animals.

IT 133081-24-0P

FL: SPN (Synthetic preparation); PREP (Preparation)  
 (prepn. and butoxylation)

RN 133081-24-0 HCAPLUS

CN 3-Pyridinecarboxylic acid, 6-hydrazino- (9CI) (CA INDEX NAME)

H2N NH N

CO2H

L4 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1991:164011 HCAPLUS

DOCUMENT NUMBER: 114:164011

TITLE: Preparation of succinimide hydrazinoarylcarboxylates and analogs as conjugating agents for biological macromolecules

INVENTOR(S): Schwartz, David A.; Abrams, Michael J.; Giandomenico, Christen M.; Subieta, Jan A.

PATENT ASSIGNEE(S): Johnson Matthey PLC, UK

SOURCE: Eur. Pat. Appl., 25 pp.

CODEN: EPHNDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY APP. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 384769	A2	19900819	EP 1990-301949	19900819
EP 384769	A2	19911127		
EP 384769	B1	19960424		
E: AT, BE, CH, DE, DK, ES, FF, GB, GR, IT, LI, LU, NL				
ZA 9001283	A	19910317	ZA 1990-1133	19900317
NO 9000838	A	19900827	NO 1990-838	19900827
NO 178186	B	19951030		
NO 178186	C	19960207		
AU 9050074	A1	19960913	AU 1990-30074	19900913
AU 630668	B2	19911105		
CA 2010800	AA	19900824	CA 1990-2010800	19900824
CA 2010800	C	20010116		
HU 53600	A1	19901128	HU 1990-970	19900128
HU 53600	B	19930519		
JP 91017356	A2	19910205	JP 1990-41389	19900205
JP 9003987	B2	20000807		
FI 95907	B	19951029	FI 1990-948	19900129
FI 95907	C	19960410		
AT 137219	E	19960515	AT 1990-301949	19900515
ES 1885890	T3	19960616	ES 1990-301949	19900616
US 5206370	A	19930427	US 1992-384641	19920520
US 5420285	A	19950530	US 1993-26426	19930530
US 5753620	A	19980219	US 1995-384641	19950219
US 6213345	B1	20010417	US 1997-384641	19970417
PRIORITY APPLN. INFO.:				
			US 1989-315170	A 19890114
			US 1990-383101	B1 19900121
			US 1992-388182	A3 19920526
			US 1993-26426	A3 19930530
			US 1995-384641	A3 19950219

OTHER SOURCE(S): MAFPAT 114:164011

BI

Q1= GE GE Q2= GE N  
A E S

AB F3DNRNH.HX, R3INRN:CR1R2, and R4NRN:CR1R2 (D = bond, CH2, CO, CSNH; R, R1, R2 = H, alkyl; R3 = aryl group Q1; A, B = CH, N; E = CO; G = group readily replaced by a primary amine; EG = maleimido; R4 = thiazolyl group Q2; X = anion) were prepa. Thus, 4-(HOLC)C6H4NHNH2 was N-protected and the product condensed with N-hydroxysuccinimide to give, after deprotection 4-(F5OLC)C6H4NHNH2.HCl (F5 = succinimido) which was conjugated with IgG and the product labeled with 99mTc. The latter gave infected/normal muscle distribution ratio of 0.3 when injected into rats having a hind leg abscess.

IT 133081-24-0P

PL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(prepn. and reaction of, in prepn. of conjugating agents for biol. macromols.)

RN 133081-24-0 HCAPLUS

CN 3-Pyridine-carboxylic acid, 6-hydrazino- (9CI) (CA INDEX NAME)

H2N NH N

CO2H

IT 6066-82-6, N-Hydroxysuccinimide

PL: RCT (Reactant); RACT (Reactant or reagent)  
(reaction of, in prepn. of conjugating agents for biol. macromols.)

RN 6066-82-6 HCAPLUS

CN 2,5-Pyrrolidinedione, 1-hydroxy- (9CI) (CA INDEX NAME)

OH

O N O

=&gt; d ilib aks hitstr 19 1-??

L9 ANSWER 1 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:67134 HCAPLUS

DOCUMENT NUMBER: 137:2655

TITLE: Attachment of benzaldehyde-modified  
oligodeoxynucleotide probes to semicarbazide-coated  
glassAUTHOR(S): Potymiragin, Mikhail A.; Lukhtanov, Eugeny A.; Reed,  
Michael W.

CORPORATE SOURCE: Epoch Biosciences, Bothell, WA, 98021, USA

SOURCE: Nucleic Acids Research (2001), 29(24), 5090-5098

CODEN: NARHAD; ISSN: 0305-1048

PUBLISHER: Oxford University Press

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Attachment of oligodeoxynucleotides (ODNs) contg. benzaldehyde (BAL) groups to semicarbazide-coated glass (SC-glass) slides is described. 5'-BAL-ODNs are prepd. using automated DNA synthesis and an acetal-protected BAL phosphoramidite reagent. The hydrophobic protecting group simplifies purifn. of BAL-ODNs by reverse phase HPLC and is easily removed using std. acid treatment. The electrophilic BAL-ODNs are stable in soln., but react specifically with semicarbazide groups to give semicarbazide silane to give SC-glass. BAL-ODNs are coupled to the SC-glass surface by a simple one-step procedure that allows rapid, efficient and stable attachment. Hand-spotted arrays of BAL-ODNs were prepd. to evaluate loading d. and hybridization properties of **immobilized** probes. Hybridization to radiolabeled target strands shows that at least 30% of the coupled ODNs were available for hybridization at max. **immobilization** d. The array was used to probe single nucleotide polymorphisms in synthetic EMA targets, and PCR products were correctly genotyped using the same macroarray. Application of this chem. to manuf. of DNA microarrays for sequence anal. is discussed.

IT 106868-88-6P

EL: ECT (Reactant); SFN (Synthetic preparation); PREP (Preparation); FACT  
(Reactant or reagent)(attachment of benzaldehyde-modified oligodeoxynucleotide probes to  
semicarbazide-coated glass)

EN 106868-88-6 HCAPLUS

CN Hydrazinecarboxamide, N-[3-(triethoxysilyl)propyl]- (9CI) (CA INDEX NAME)

OEt O

EtO Si (CH<sub>2</sub>)<sub>3</sub> NH C NH NH<sub>2</sub>

OEt

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 2 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:843748 HCAPLUS

DOCUMENT NUMBER: 135:37103

TITLE: Water dispersion compositions useful as coatings of  
metals especially automobilesINVENTOR(S): Yamauchi, Tetsuaki; Takahashi, Hiroaki; Takada,  
Toshihiko

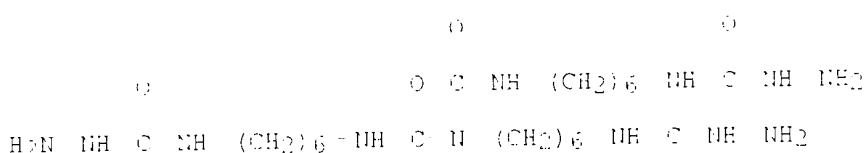
PATENT ASSIGNEE(S): Asahi Chemical Industry Co., Ltd., Japan  
 SOURCE: Jpn. Kokai Tokkyo Koho, 25 pp.  
 CODEN: JKKXAP  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY APP. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2001333123	A1	20011120	JP 2000-139946	20000512
PRIORITY APPLN. INFO.:			JP 2000-139946	20000512

AB Title compns. comprise (i) carbonyl group-contg. aq. film forming resins comprising aq. polycarbonyl compds. and (ii) thermal **crosslinking** agents. The compns. are useful as intermediate and/or top base coatings for automobiles, esp. as three coat-one bake, and give coating films with good appearance. Thus, (a) an intermediate water resistant coating compn., (b) a white top base water resistant coating compn., both comprising aq. polycarbonyl compd. obtained from Bu acrylate, diacetone acrylamide, 2-hydroxyethyl methacrylate, methacrylic acid, Me methacrylate, styrene, 2,2'-azobis(2,4-dimethylvaleronitrile), and N,N-dimethylethanolamine, aq. film forming resin obtained from Bu acrylate, diacetone acrylamide, 2-hydroxyethyl methacrylate, Latemul S 150A, methacrylic acid, Me methacrylate, trimethylolpropane triacrylate, N,N-dimethylethanolamine, and ammonia, and Cymel 154, and (c) an acrylic clear coating compn. were applied on an electrodeposited coated plate (wet on wet method), electrostatically coated, and baked at 150.degree. for 25 min to give a coating film with good appearance.

IT **175870-12-9P**  
 FL: IMF (Industrial manufacture); MOA (Modifier or additive use); PREP (Preparation); USES (Uses)  
 (water dispersion intermediate and/or top base coating compns. giving cured coating films with good appearance)

RN 175870-12-9 HCAPLUS  
 CN 1,9,11,13,20-Pentaazaheneicosanedioic acid, 11-[6-[(hydrazinocarbonyl)amino]hexyl]-10,12-dioxo-, dihydrazide (9CI) (CA INDEX NAME)



IT **374620-65-2P 374620-67-4P 374620-76-5P**  
 FL: IMF (Industrial manufacture); TEM (Technical or engineered material use); PREP (Preparation); USES (Uses)  
 (water dispersion intermediate and/or top base coating compns. giving cured coating films with good appearance)

RN 374620-65-2 HCAPLUS  
 CN 1,9,11,13,20-Pentaazaheneicosanedioic acid, 11-[6-[(hydrazinocarbonyl)amino]hexyl]-10,12-dioxo-, dihydrazide, polymer with butyl 2-propenoate, N-(1,1-dimethyl-3-oxobutyl)-2-propanamide, ethenylbenzene, 2-ethyl-2-[[[(1-oxo-2-propenyl)oxy]methyl]-1,3-propanediyl di-2-propenoate, formaldehyde, 2-hydroxyethyl 2-methyl-2-propenoate, Latemul S 150A, methyl 2-methyl-2-propenoate, 2-methyl-2-propenoic acid

and 1,3,5-triazine-2,4,6-triamine, ammonium salt, compd. with  
2-(dimethylamino)ethanol (9CI) (CA INDEX NAME)

CM 1

CFN 113-01-0  
CMF C4 H11 N O

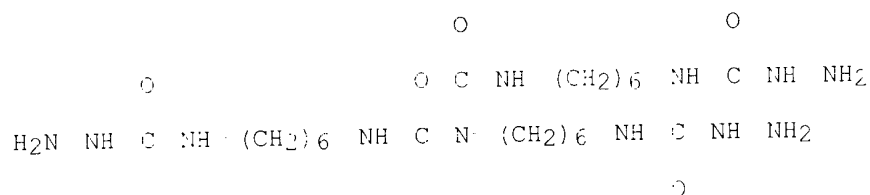
Me2N CH2 CH2 OH

CM 2

CFN 374620-64-1  
CMF (C13 H50 N12 O5 . C15 H20 O6 . C9 H15 N O2 . C8 H8 . C7 H12 O2 . C6  
H10 O3 . C5 H8 O2 . C4 H6 O2 . C3 H6 N6 . C H2 O . Unspecified)x  
CCI EMS

CM 3

CFN 175570-12-9  
CMF C23 H50 N12 O5



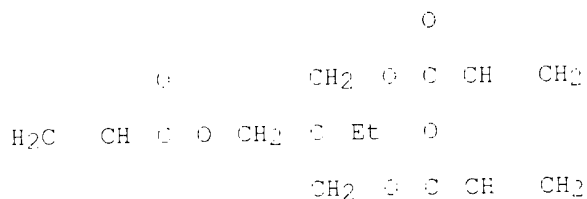
CM 4

CFN 113255-53-1  
CMF Unspecified  
CCI MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 5

CFN 15625-89-5  
CMF C15 H20 O6





CM 6

CRN 2873-97-4  
CMF C9 H15 N O2

O

H2C CH C NH C

Me C CH2 C Me

Me

CM 7

CRN 268-77-9  
CMF C6 H10 O3

H2C O

Me C C O CH2 CH2 OH

CM 8

CRN 141-30-2  
CMF C7 H12 O2

O

n-BuO C CH CH2

CM 9

CRN 108-78-1  
CMF C3 H6 N6

NH2

N H

H2N N NH2

CM 10

CRN 100-42-5  
CMF C8 H8

H<sub>2</sub>C CH Ph

CM 11

CFN 80-62-6  
CMF C5 H8 O2

H<sub>2</sub>C O

Me C O OMe

CM 12

CFN 79-41-4  
CMF C4 H6 O2

CH<sub>2</sub>

Me C CO<sub>2</sub>H

CM 13

CFN 50-00-0  
CMF C H2 O

H<sub>2</sub>C O

EN 574620-67-4 HCAPLUS

CN 2,9,11,13,20-Pentaazaheneicosanedioic acid, 11-[6-(hydrazinocarbonyl)amino]hexyl]-10,12-dioxo-, dihydrazide, polymer with butyl 2-propenoate, N-(1,1-dimethyl-3-oxobutyl)-2-propenamide, ethenylbenzene, formaldehyde, 2-hydroxyethyl 2-methyl-2-propenoate, Latemul S 180A, methyl 2-methyl-2-propenoate, 2-methyl-2-propenoic acid and 1,3,5-triazine-2,4,6-triamine, ammonium salt, compd. with N-(dimethylamino)ethanol (9CI) (CA INDEX NAME)

CM 1

CFN 108-01-0  
CMF C4 H11 N O

Me<sub>2</sub>N CH<sub>2</sub> CH<sub>2</sub> OH

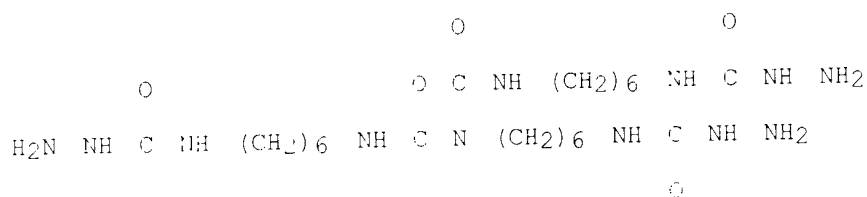
CM 2

CFN 574620-66-3

CMF (C23 H50 N12 O5 . C9 H15 N O2 . C8 H8 . C7 H12 O2 . C6 H10 O3 . C5 H8  
O . C4 H6 O2 . C3 H6 N6 . C H2 O . Unspecified)x  
CCI FMS

CM 3

CFN 175370-12-9  
CMF C23 H50 N12 O5



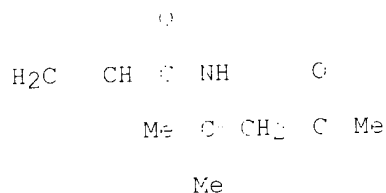
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CFN 111251-51-1  
CMF Unspecified  
CCI MAN

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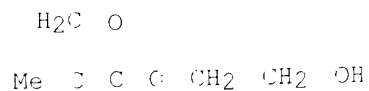
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CFN 2873-97-4  
CMF C9 H15 N O2



CM 6

CFN 363-77-3  
CMF C6 H10 O3



CM 7

CFN 141-32-2  
CMF C7 H12 O2

O

n-BuO C CH CH<sub>2</sub>

CM 8

CRN 108-78-1  
CMF C3 H6 N6

NH<sub>2</sub>

N N

H<sub>2</sub>N N NH<sub>2</sub>

CM 9

CRN 100-42-5  
CMF C8 H8

H<sub>2</sub>C CH Ph

CM 10

CRN 80-62-6  
CMF C5 H8 O2

H<sub>2</sub>C C

Me C C OMe

CM 11

CRN 79-41-4  
CMF C4 H6 O2

CH<sub>2</sub>

Me C CO<sub>2</sub>H

CM 12

CRN 50-00-0

CMF C H2 O

H2C O

RN 374620-76-5 HCAPLUS  
CN 2,3,11,13,20-Pentaazaheneicosanedioic acid, 11-[6-  
[(hydrazinocarbonyl)amino]hexyl]-10,12-dioxo-, dihydrazide, polymer with  
butyl 2-propenoate, cyclohexyl 2-methyl-2-propenoate, N-(1,1-dimethyl-3-  
oxobutyl)-2-propenamide, formaldehyde, 2-hydroxyethyl 2-methyl-2-  
propenoate, Latemul S 180A, methyl 2-methyl-3-propenoate,  
2-methyl-2-propenoic acid and 1,3,5-triazine-2,4,6-triamine, ammonium  
salt, compd. with 2-(dimethylamino)ethanol (PCI) (CA INDEX NAME)

CM 1

CFN 105-01-0  
CMF C4 H11 N O

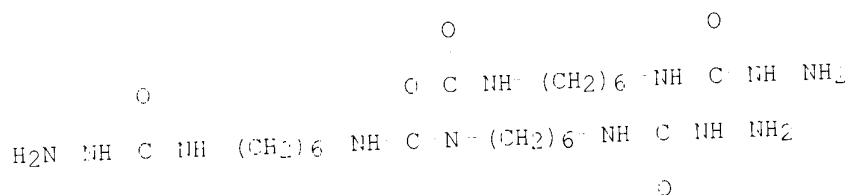
Me2N CH2 CH2 OH

CM 2

CFN 374620-76-4  
CMF (C12 H50 N12 O5 . C10 H16 O2 . C9 H15 N O2 . C7 H12 O2 . C6 H10 O3 .  
C5 H8 O2 . C4 H6 O2 . C3 H6 N6 . C H2 O . Unspecified)x  
CCI PMS

CM 3

CFN 105-70-12-9  
CMF C13 H50 N12 O5



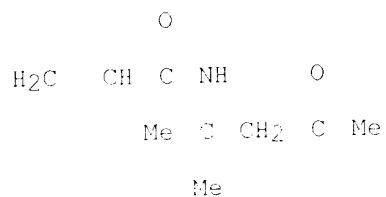
CM 4

CFN 113255-53-1  
CMF Unspecified  
CCI MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

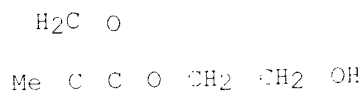
CM 5

CFN 2373-97-4  
CMF C9 H15 N O2



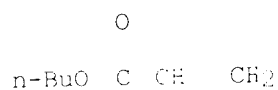
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CFN 863-77-9  
CMF C6 H10 O3



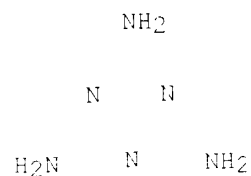
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CFN 141-32-2  
CMF C7 H12 O2



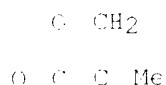
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CMF C3 H6 N6



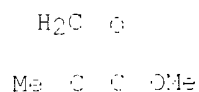
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CFN 101-43-9  
CMF C10 H16 O2



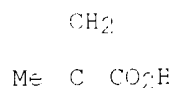
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CPN 30-62-6  
CMF C5 H8 O2



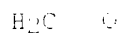
CM 11

CPN 79-41-4  
CMF C4 H6 O2



CM 12

CPN 50-00-0  
CMF C H2 O



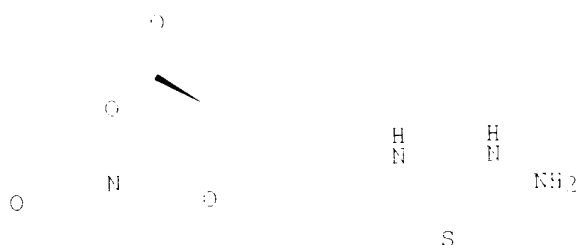
L9 ANSWER 3 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2001:713305 HCAPLUS  
DOCUMENT NUMBER: 135:272864  
TITLE: Hydrazine-based and carbonyl-based  
**bifunctional crosslinking** reagents  
for biomolecules, drugs, and synthetic polymers  
INVENTOR(S): Schwartz, David A.  
PATENT ASSIGNEE(S): Solulink, Inc., USA  
SOURCE: PCT Int. Appl., 97 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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 WO 2001070685 A2 20010927 WO 2001-US9252 20010322  
 WO 2001070685 A3 20020327  
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 GR, HC, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LF, LR, LS,  
 LT, LU, LV, MA, MD, ME, MG, MW, MX, MY, NZ, NG, NI, NO, PL, PT, RO,  
 RU, SD, SE, SG, SI, SK, SL, SN, ST, SV, TH, TJ, TR, UA, UG, US, UZ,  
 VN, YU, ZA, ZW, AM, AN, BT, BG, BE, ME, FI, TJ, TM  
 RW: GH, GM, KE, LS, MW, NG, SD, SE, SG, TH, TN, TZ, UA, BE, CH, CY,  
 IE, UK, ES, FI, FR, GB, GR, IE, IL, LU, MC, NL, PT, SE, TR, BF,  
 BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
 US 2002013857 A1 20030116 US 2001-513978 20010312  
 EP 1314693 A2 20030604 EP 2001-920666 20010312  
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, IL, LU, NL, SE, MC, PT,  
 IE, SI, LT, LV, FI, FO, MK, CY, AL, TR  
 US 2000-191134P 20000323  
 WO 2001-US9252 W 20010322  
 PRIORITY APPLN. INFO.:  
 OTHER SEARCH(S): MARPAT 195:072864  
 AB Reagents and methods are provided for **bifunctional crosslinking and immobilizing** biomols., drugs, and synthetic polymers. The reagents of formula  $BPACHNEA.bol.HX$  [wherein A =  $NHCH_2$ ,  $NHCH_2$ ,  $NHNECH_2$ ,  $NHNECH_2$ , or a direct bond; B = an amino or thio reactive moiety; F = specified aliph. divalent groups exclg. any combination of cycloalkylene,  $C(F10)2$ ,  $CF10:CF10$ ,  $C:CF12R13$ ,  $CF12R13$ ,  $C:tpbond.C$ ,  $O$ ,  $SCa$ ,  $NF10$ ,  $N-R12R13$ ,  $Cl$ , etc.; a = 0-2; x = 0-3; G = G or  $NF10$ ; L = S, O, or  $NH10$ ; R10 = specified monovalent groups; R12 and R13 = independently H, (cyclo)alkyl, alkenyl, alkynyl, or (hetero)aryl; or R12 and R13 together form cycloalkylene or alkenylene; X = neg. counterion; or a deriv. thereof] possess a thiol or amino reactive group and a hydrazine or oxyamino moiety. **Conjugates and immobilized biomols.** are also provided. For example, hydrazinonicotinic acid was converted to the acetone hydrazone and treated with N-hydroxysuccinimide to give the **crosslinking agent**, succinimide 6-hydrazinonicotinate acetone hydrazone (I), in 43% yield. A soln. of ovalbumin in PBS and BETA was added to a soln. of I in DMF and the mixt. incubated at room temp. for 4 h to afford the hydrazine-modified protein, which exhibited a molar extinction coeff. of 10,000 at 360 nm.  
 IT **362522-51-8P**  
 RL: RCT (Reactant); SPE (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
 (crosslinking agent; prepn. of hydrazine- and carbonyl-based bifunctional crosslinking agents and use with biomols., drugs, and synthetic polymers)  
 RN 362522-51-8 HCAPLUS  
 CN Hydrazinecarbothioamide, N-[[trans-4-[[[2,5-dioxo-1-pyrrolidinyl]oxy]carbonyl]cyclohexyl]methyl]-, monohydrochloride (90I)  
 (CA INDEX NAME)

Relative stereochemistry.





L9 ANSWER 4 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2001:247326 HCAPLUS  
DOCUMENT NUMBER: 134:42166  
TITLE: Preparation of a coating, a coated substrate, an  
adhesive, a film or sheet, and the coating mixture to  
be used  
INVENTOR(S): Hesselmans, Laurentius Cornelius Josephus; Spek, Dirk  
Pieter  
PATENT ASSIGNEE(S): Stahl International B.V., Neth.  
SOURCE: PCT Int. Appl., 33 pp.  
CODEN: PIKMD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 100102451	A2	20010405	WO 1000-NL699	20000929
WO 100102451	A3	20011025		
W:	AE, AL, AM, AT, AU, AZ, BA, BB, BE, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, GR, GU, ID, IL, IN, IS, JP, KE, KG, KF, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, ML, ME, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, VC, VN, YU, ZA, ZW, AM, AZ, BY, CG, CH, CI, CM, CN, CO, CR, CU, CY, EE, EG, ES, FI, FR, GB, GR, GU, HK, HN, HU, IE, IL, IN, IS, IT, JP, KE, KG, KH, KI, KP, KR, KZ, LA, LB, LC, LI, LU, LT, LV, LY, MA, MD, ME, MG, MK, MN, MO, MP, MQ, MR, MT, MU, MV, MW, MX, MY, MZ, NA, NC, NE, NG, NI, NL, NO, NP, NR, NU, NZ, OI, OM, OS, PA, PE, PG, PH, PK, PL, PM, PN, PR, PT, PU, PY, QA, QG, QH, QI, QJ, QK, QL, QM, QN, QO, QP, QQ, QR, QS, QT, QU, QV, QW, QX, QY, QZ, RA, RB, RC, RD, RE, RF, RG, RH, RI, RJ, RK, RL, RM, RN, RO, RP, RR, RS, RT, RU, RV, RW, RX, RY, RZ, SA, SB, SC, SD, SE, SF, SG, SH, SI, SJ, SK, SL, SM, SN, SO, SP, SR, SS, ST, SU, SV, SW, SX, SY, SZ, TA, TB, TC, TD, TE, TF, TG, TH, TI, TJ, TK, TL, TM, TN, TO, TP, TR, TS, TT, TV, TW, TX, TY, TZ, UA, UB, UC, UD, UE, UF, UG, UH, UI, UJ, UK, UL, UM, UN, UO, UP, UQ, UR, US, UT, UV, UW, UX, UY, UZ, VA, VB, VC, VD, VE, VF, VG, VH, VI, VJ, VK, VL, VM, VN, VO, VP, VQ, VR, VS, VT, VU, VV, VW, VX, VY, VZ, WA, WB, WC, WD, WE, WF, WG, WH, WI, WJ, WK, WL, WM, WN, WO, WP, WQ, WR, WS, WT, WU, WV, WX, WY, WZ, XA, XB, XC, XD, XE, XF, XG, XH, XI, XJ, XK, XL, XM, XN, XO, XP, XQ, XR, XS, XT, XU, XV, XW, XX, XY, XZ, YA, YB, YC, YD, YE, YF, YG, YH, YI, YJ, YK, YL, YM, YN, YO, YP, YQ, YR, YS, YT, YU, YV, YW, YY, YZ, ZA, ZB, ZC, ZD, ZE, ZF, ZG, ZH, ZI, ZJ, ZK, ZL, ZM, ZN, ZO, ZP, ZQ, ZR, ZS, ZT, ZU, ZV, ZW, ZX, ZY, ZZ			
FW:	GH, GM, KE, LS, MW, MC, SD, SL, SO, TZ, UG, JW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, CA, CN, GW, ML, MP, NE, SN, TD, TG			
NL 1013179	A2	20010402	NL 1999-1013179	19990930
EE 1033991	A2	20000818	EE 2000-970320	20000929
E:	AE, BE, CH, DE, DK, ES, FF, GB, GE, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, FO, MK, CY, AL			
BE 2000014669	A	20021001	BE 2000-14669	20000929
JP 2003510431	T2	20030518	JP 2001-526598	20000929
PRIORITY APPLN. INFO.:			NL 1999-1013179 A	19990930
			WO 2000-NL699 W	20000929

Searched by Mary Jane Fuhl 605-1154

conditions, is applied onto a substrate at ambient temp., followed by heating. At ambient temp. the compd. contg. reactive E is a solid material, a powder, a granule, a flake or grind or a ground mixt. The coatings, coated substrates, adhesives, films, sheets, impregnated substrates, synthetic leathers, in-mold coatings, coated leathers, coated poly(vinyl chloride), coated nonwovens, coated coagulated polyurethane substrates, breathable coated substrates, are obtained by applying the the title process.

IT 332421-29-1P 332421-30-4P

FI: IMF (Industrial manufacture); PRP (Properties); TEM (Technical or engineered material use); PREP (Preparation); USES (Uses)  
(coating or film; for coating, adhesive, a film or sheet formulated from a reactive mixt. of long pot life, fast reaction, and low toxic vapors)

RN 332421-29-1 HCAPLUS

CN Hexanedioic acid, polymer with 2,2-dimethyl-1,3-propanediol, 2-ethyl-1-(hydroxymethyl)-1,3-propanediol, 1,6-hexanediol, N,N'-1,6-hexanediylbis[hydrazinecarboxamide] and 5-isocyanato-1-(isocyanatomethyl)-1,3,3-trimethylcyclohexane (9C1) (CA INDEX NAME)

CM 1

CRN 51440-79-1  
CMF C3 H20 N6 O2

O

O

H2N NH C NH (CH2)5 NH C NH NH2

CM 2

CRN 4083-71-9  
CMF C13 H16 N2 O2

OCN

Me

CH1 NCO

Me Me

CM 3

CRN 629-11-5  
CMF C6 H14 O2

HO (CH2)6 OH

CM 4

CPN 126-30-7  
CMF C6 H12 O2

Me

HO CH<sub>2</sub> C CH<sub>2</sub> OH

Me

CM 5

CPN 124-04-9  
CMF C6 H10 O4

HO<sub>2</sub>C (CH<sub>2</sub>)<sub>4</sub> CO<sub>2</sub>H

CM 6

CPN 77-89-6  
CMF C6 H14 O3

CH<sub>2</sub> OH

HO CH<sub>2</sub> C Et

CH<sub>2</sub> OH

RM 330421-30-4 HCAPLUS  
CN Hexanediolic acid, polymer with 2,2-dimethyl-1,3-propanediol,  
2-ethyl-2-(hydroxymethyl)-1,3-propanediol, 1,6-hexanediol,  
N-[3-[[[(hydrazinocarbonyl)amino]methyl]-3,5,5-  
trimethylcyclohexyl]hydrazinecarboxamide and 5-isocyanato-1-  
isocyanatomethyl-1,3,3-trimethylcyclohexane (9CI) (CA INDEX NAME)

CM 1

CPN 52384-45-4  
CMF C12 H26 N6 O2

O

H<sub>2</sub>N CH C NH

Me

O

CH<sub>2</sub> NH C NH NH<sub>2</sub>

Me Me

CM 2

CRN 4098-71-9  
CMF C11 H18 N2 O2

OCN Me  
CH2 NCO

Me Me

CM 3

CRN 629-11-8  
CMF C6 H14 O2

HO (CH2)6 OH

CM 4

CRN 116-30-7  
CMF C5 H12 O2

Me

HO CH2 C CH2 OH

Me

CM 5

CRN 114-04-9  
CMF C6 H10 O4

HO2C (CH2)4 CO2H

CM 6

CRN 77-99-6  
CMF C6 H14 O3

$$\begin{array}{c} \text{CH}_2 \quad \text{OH} \\ | \\ \text{HC} \quad \text{CH}_2 \quad \text{C} \quad \text{Et} \end{array}$$

$$\begin{array}{c} \text{CH}_2 \quad \text{OH} \\ | \\ \text{HC} \quad \text{CH}_2 \quad \text{C} \quad \text{Et} \end{array}$$

IT 51440-70-1P 52284-45-4P

FL: IMF (Industrial manufacture); FCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)

(curative; for coating, adhesive, a film or sheet formulated from a reactive mixt. of long pot life, fast reaction, and low toxic vapors)

RN 51440-70-1 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,6-hexanediylbis- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{O} \quad \quad \quad \text{O} \\ | \quad \quad \quad | \\ \text{H}_2\text{N} \quad \text{NH} \quad \text{C} \quad \text{NH} \quad (\text{CH}_2)_6 \quad \text{NH} \quad \text{C} \quad \text{NH} \quad \text{NH}_2 \end{array}$$

H<sub>2</sub>N NH C NH (CH<sub>2</sub>)<sub>6</sub> NH C NH NH<sub>2</sub>

RN 52284-45-4 HCAPLUS

CN Hydrazinecarboxamide, N-[3-[[[(hydrazinocarbonyl)amino]methyl]-3,5,5-trimethylcyclohexyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{O} \quad \quad \quad \text{Me} \quad \quad \quad \text{O} \\ | \quad \quad \quad | \quad \quad \quad | \\ \text{H}_2\text{N} \quad \text{NH} \quad \text{C} \quad \text{NH} \quad \quad \quad \text{CH}_2 \quad \text{NH} \quad \text{C} \quad \text{NH} \quad \text{NH}_2 \end{array}$$

H<sub>2</sub>N NH C NH

CH<sub>2</sub> NH C NH NH<sub>2</sub>

Me Me

IT 32251-26-6 126953-51-3 332421-34-8

FL: TEM (Technical or engineered material use); USES (Uses)

(curative; for coating, adhesive, a film or sheet formulated from a reactive mixt. of long pot life, fast reaction, and low toxic vapors)

RN 32251-26-6 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,2-ethanediylbis- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{O} \quad \quad \quad \text{O} \\ | \quad \quad \quad | \\ \text{H}_2\text{N} \quad \text{NH} \quad \text{C} \quad \text{NH} \quad \text{CH}_2 \quad \text{CH}_2 \quad \text{NH} \quad \text{C} \quad \text{NH} \quad \text{NH}_2 \end{array}$$

H<sub>2</sub>N NH C NH CH<sub>2</sub> CH<sub>2</sub> NH C NH NH<sub>2</sub>

RN 126953-51-3 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,3-propanediylbis- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{O} \quad \quad \quad \text{O} \\ | \quad \quad \quad | \\ \text{H}_2\text{N} \quad \text{NH} \quad \text{C} \quad \text{NH} \quad (\text{CH}_2)_3 \quad \text{NH} \quad \text{C} \quad \text{NH} \quad \text{NH}_2 \end{array}$$

H<sub>2</sub>N NH C NH (CH<sub>2</sub>)<sub>3</sub> NH C NH NH<sub>2</sub>

RN 332421-34-8 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,4-butanediylbis- (9CI) (CA INDEX NAME)

H2N NH C NH (CH2)4 NH C NH NH2

L9 ANSWER 5 OF 18 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:167451 HCAPLUS

DOCUMENT NUMBER: 134:198050

TITLE: Radiopharmaceutical products and their preparation procedure

INVENTOR(S): Bellande, Emmanuel; Jallet, Pierre; Teninot, Benoit

PATENT ASSIGNEE(S): Cis Bio International, Fr.

SOURCE: PCT Int. Appl., 46 pp.

CODEN: PIXMD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY APP. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 1991015746	A1	20010308	WO 2000-IB1161	20000923
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BR, BG, BF, BY, BE, CA, CH, CN, CR, CU, CE, DE, DK, DM, DO, EE, ES, FI, GE, GD, GE, GH, GM, GR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LF, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, ME, MC, MG, NE, NL, NO, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, VE, VN, YU, ZA, ZW, AM, AE, BY, BG, EL, ES, FR, FI, TM			
FW:	CH, CM, KE, LS, MW, ME, SD, SL, SE, TD, UG, ZW, AT, PE, CH, CY, IE, DK, ES, FI, FR, GE, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, IF, IG, OL, OM, SA, SN, SW, ML, MF, NE, SE, TD, TG			
EP 2747769	A1	20010308	EP 1999-10970	19990601
EP 2000013729	A	20010507	EP 2000-11729	20000623
EP 1210137	A1	20010605	EP 2000-951784	20000823
P:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, FO, MK, CY, AL			
JP 200308495	T2	20030304	JP 2001-520157	20030623
EE 200309195	A	20030415	EE 2001-105	20030623
BZ 200433	A	20040430	BZ 2003-296438	20040623
NO 2002001001	A	20020411	NO 2003-1901	20030623
PRIORITY APPLN. INFO.:			EP 1999-10970 A 19990601	
			WO 2000-IB1161 W 20000923	

OTHER SOURCE(S): MAFPAT 134:198050

AB The present invention relates to radiopharmaceutical products and their prepn. procedure. These products can be used for pulmonary scintigraphy or for therapy. They comprise a polysaccharide and sequestering groups of formulas R-NH-, R-N=, and R-N(R')N= in which R is a hydrocarbon or arom. group comprising at least one atom of sulfur, and R' is an atom of hydrogen or an alkyl grouping such as Me, said sequestering groups forming a chelate type complex with a radioactive metal such as technetium.

IT 3766-55-ODP, 4-Allyl 3-thiosemicarbazide, radiolabeled reaction product with oxidized starch 6610-29-3DP, 4-Methyl 3-thiosemicarbazide, radiolabeled reaction product with oxidized starch FI: BPR (Biological process); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); FIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)

(radiopharmaceutical kits for scintigraphy)  
 RN 0766-55-0 HCAPLUS  
 CN Hydrazinecarbothioamide, N-2-propenyl- (9CI) (CA INDEX NAME)

S

H2N NH C NH CH2 CH CH2

RN 6610-29-2 HCAPLUS  
 CN Hydrazinecarbothioamide, N-methyl- (9CI) (CA INDEX NAME)

S

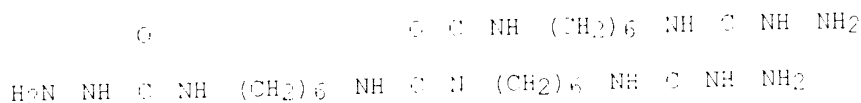
MeNH C NH NH2

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS  
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 6 OF 25 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1999:114037 HCAPLUS  
 DOCUMENT NUMBER: 130:200010  
 TITLE: lightweight cellular concrete having waterproof  
 coatings and its preparation  
 INVENTOR(S): Ito, Yasuyuki; Watanabe, Tomiyo; Nakanishi, Masuhiko  
 PATENT ASSIGNER(S): Asahi Chemical Industry Co., Ltd., Japan  
 SOURCE: Jpn. Kokai Tokkyo Koho, 3 pp.  
 CODEN: JKEMAF  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 11043385	A2	19990216	JP 1997-198765	19970724
			JP 1997-198765	19970724

PRIORITY APPLN. INFO.:  
 AB The prepn. involves the following steps; (1) impregnating lightweight  
 cellular concrete with an aq. soln. contg. a hardenable resin which shows  
 water soly. before cross linking, (2) **crosslinking** the resin,  
 and (3) forming a coating on the surface. The aq. soln. may contain a  
 hardening agent. The resulting concrete products are also claimed.  
 IT **175870-12-9P**  
 EL: INF (Industrial manufacture); MDA (Modifier or additive use); RCT  
 (Reactant); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (crosslinking agent; prepn. of lightweight cellular concrete  
 having **crosslinked** polymer layers and waterproofing coating  
 layers)  
 RN 175870-12-9 HCAPLUS  
 CN 2,9,11,13,20-Pentaazaheneicosanedioic acid, 11-[6-  
 [(hydrazinocarbonyl)amino]hexyl]-19,12-dioxo-, dihydrazide (9CI) (CA  
 INDEX NAME)



LA ANSWER 7 OF 39 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 130:724187 HCAPLUS

DOCUMENT NUMBER: 130:14992

TITLE: Isophoronebis(semicarbazides), their preparation, their semicarbazones, and room-temperature-curable water-resistant coating compositions with good storage stability containing them

INVENTOR(S): Yokota, Masahisa; Miyazaki, Takayuki; Ueyanagi, Kaoru

PATENT ASSIGNEE(S): Asahi Chemical Industry Co., Ltd., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 11 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

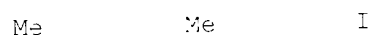
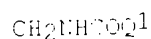
LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 10298158	A2	19981110	JP 1997-120103	19970424
PRIORITY APPLN. INFO.:			JP 1997-120103	19970424
OTHER SOURCE(S):		MARPAT 130:14992		

SI



AB The semicarbazides I (Q1, Q2 = NR3NH2 (R3 = H, C1-20 alkyl, alicyclic group, aryl), NHNR3F4NR3NH2 (R4 = linear or branched C2-20 alkylene, C5-20 cycloalkylene, C6-10 arylene which may be substituted with C1-8 alkyl or alkoxy), NHNR3COR4CONR3NH2, NHNR3CONHNH)xCONHNH2 (x = 1-5), NHNR3CONHNH4NHCONR3NH2) are prep'd. by treatment of isophorone diisocyanate and hydrazines. Semicarbazones are prep'd. by treatment of I with R1R2CO (R1, R2 = H, linear or branched C2-20 aliph. group, C5-20 alicyclic group, (un)substituted aryl; R1 and R2 may be bonded to each other forming a ring). The coating compns. contain (A) I, and/or (B) the above semicarbazones, and (C) polycarbonyl compds. at (C)/[(A)+(B)] = 99.9:0.1-10/90. The compns. provide coating having high hardness and good waterproofness. An aq. emulsion contained methacrylic acid-Me methacrylate-Bu acrylate-diisobutyl acrylamide copolymer and isophorone bis(semicarbazide).

IT 216143-35-0P

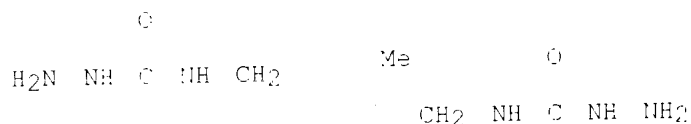
RL: IMF (Industrial manufacture); MDA (Modifier or additive use); RCT



(Reactant); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (prepn. of isophoronebis(semicarbazides) as **crosslinking**  
 agents for room-temp.-curable coating compns. for high hardness and  
 good waterproofness)

RN 216143-35-0 HCAPLUS

CN Hydrazinecarboxamide, N,N'-[ (1,5,5-trimethyl-1,3-  
 cyclohexanediyl)bis(methylene)]bis- (9CI) (CA INDEX NAME)



Me Me

IT 216143-36-1P

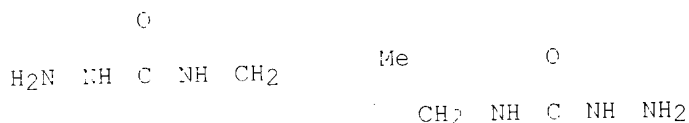
FI: IMF (Industrial manufacture); PRP (Properties); TEM (Technical or  
 engineered material use); PREP (Preparation); USES (Uses)  
 (prepn. of isophoronebis(semicarbazides) as **crosslinking**  
 agents for room-temp.-curable coating compns. for high hardness and  
 good waterproofness)

RN 216143-36-1 HCAPLUS

CN 2-Propenoic acid, 2-methyl-, polymer with butyl 2-propenoate,  
 N-(1,1-dimethyl-3-oxobutyl)-2-propenamide, methyl 2-methyl-2-propenoate  
 and N,N'-[ (1,5,5-trimethyl-1,3-cyclohexanediyl)bis(methylene)]bis[hydrazin-  
 ecarboxamide] (9CI) (CA INDEX NAME)

CM 1

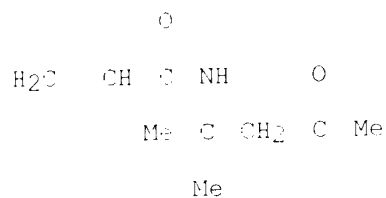
ERN 216143-35-0  
 IMF 013 H23 N6 02



Me Me

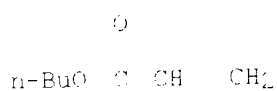
CM 2

ERN 2873-97-4  
 IMF 09 H15 N 02



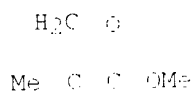
CM 3

CRN 141-32-2  
CMF C7 H12 O2



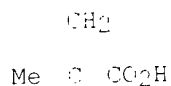
CM 4

CRN 80-62-6  
CMF C5 H8 O2



CM 5

CRN 79-41-4  
CMF C4 H6 O2



L9 ANSWER 8 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1998:112262 HCAPLUS  
 DOCUMENT NUMBER: 128:196654  
 TITLE: Polypeptides having a single covalently bound  
 N-terminal water-soluble polymer  
 INVENTOR(S): Wei, Ziping; Menon-rudolph, Sunitha; Ghosh-Dastidar,  
 Pradip  
 PATENT ASSIGNEE(S): Ortho Pharmaceutical Corp., USA  
 SOURCE: PCT Int. Appl., 51 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

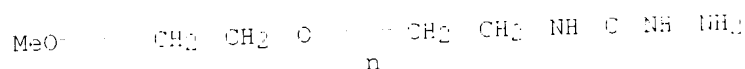
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 98 05363	A2	19970111	WO 1997-US13755	19970801
WO 98 05363	A2	19970507		
Z: AL, AM, AT, AU, AZ, BA, BB, BG, BE, BY, CA, CH, CN, CU, CZ, DE, EE, ES, FI, FR, GE, GR, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, MY, NG, NL, PT, PO, RU, SD, SE, SG, SI, SK, SL, ST, TH, TR, TT, UA, UG, UZ, VN, YU, YW				
FW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MD, NL, PT, SE				
AU 8730085	A1	19980225	AU 1997-39085	19970801
EP 9711009	A	19960817	EP 1997-11009	19970801
CN 1116176	A	19990818	CN 1997-196529	19970801
EP 984702	A2	19991212	EP 1997-936497	19970801
F: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI				
NZ 343993	A	20000113	NZ 1997-353995	19970801
JP 20000515853	T2	20001121	JP 1998-508173	19970801
RU 2199847	C2	20030217	RU 1999-103679	19970801
NO 9400465	A	19990523	NO 1999-465	19990501
MX 9401184	A	20000301	MX 1999-1184	19990301
PRIORITY ATELN. INFO.:				
US 1996-030311 P 19960902				
WO 1997-US13755 W 19970801				

AB This invention provides compns. consisting essentially of a polypeptide such as erythropoietin and a water-sol. polymer such as PEG covalently bound thereto at the N-terminal .alpha.-carbon atom via a hydrazone or reduced hydrazone bond, or an oxime or reduced oxime bond. This invention also provides methods of making the instant compns., pharmaceutical compns. comprising same, and kits for use in prepg. same.

IT 167394-62-9  
 PL: RCT (Reactant); RACT (Reactant or reagent)  
 polypeptides having a single covalently bound N-terminal water-sol. polymer)

RN 167394-62-9 HCAPLUS  
 CN Poly(oxo-1,2-ethanediyl), .alpha.-[2-[(hydrazinocarbonyl)amino]ethyl]-.omega.-methoxy- (BCI) (CA INDEX NAME)

O



L9 ANSWER 9 OF 36 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1997:390850 HCAPLUS  
 DOCUMENT NUMBER: 117:1745  
 TITLE: Reagent for the detection and isolation of carbohydrates or glycan receptors  
 INVENTOR(S): Witzele, Manfred; Fernholz, Erhard; Von Der Eltz, Herbert  
 PATENT ASSIGNEE(S): Boehringer Mannheim GmbH, Germany  
 SOURCE: Eur. Pat. Appl., 29 pp.  
 DLEN: EPXXLW  
 DOCUMENT TYPE: Patent  
 LANGUAGE: German  
 FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 709490	A1	19970423	EP 1996-116773	19961018
EP 709490	B1	20011319		
F: DE, ES, FR, GB, IT				
DE 19539008	A1	19970423	DE 1995-19539008	19951019
US 6118546	B1	20010417	US 1996-733736	19961018
JP 09176106	A2	19970703	JP 1996-277834	19961021
			DE 1995-19539008 A	19951019

PRIORITY APPLN. INFO.:

OTHER SOURCE(S):

MARPAT 127:2745

AB The finding concerns compds., which contain a chromophore and a ligand (e.g., biotin or a biotin deriv.) that can bind to streptavidin and/or avidin, that are suitable for binding to mois. that contain an aldehyde, ketone, hemiacetal, and/or hemiketal function. The finding also concerns **conjugates** formed from these compds. as well as a method for detecting or isolating carbohydrates or glycan receptors by using such **conjugates**.

IT 190126-38-6P

PL: ARG (Analytical reagent use); RCT (Reactant); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); FACT (Reactant or reagent); USES (Uses)  
(reagent for detecting and isolating carbohydrates or glycan receptors)

RN 190126-38-6 HCAFLUS

CN Hydrazinecarboxamide, N-[2-[[2-[(4-hydroxyphenyl)azo]benzoyl]amino]ethyl]-, mono(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

CM 1

CRM 190126-37-5

CMF C16 H18 N6 O3

OH

CH N

C NH CH2 CH2 NH C NH NH2

CM 2

CRM 70-06-1

CMF C12 H13 F3 O2

F

F C CO2H

F

L9 ANSWER 10 OF 38 HCAFLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1997:00002 HCAPLUS  
 DOCUMENT NUMBER: 126:200300  
 TITLE: Di- and triaminoguanidines, and methods of use  
 INVENTOR(S): Wagle, Philip R.; Ulrich, Peter J.; Jensen, Anthony  
 PATENT ASSIGNEE(S): Altec Inc., USA; Rockefeller University  
 SOURCE: U.S., 13 pp., Cont.-in-part of U.S. Ser. No. 274,243,  
 abandoned.  
 CODEN: USREAM  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 33  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5612382	A	19970318	US 1995-487089	19950607
EP 322402	A2	19890619	EP 1989-101406	19890319
EP 322402	A2	19891025		
EP 322402	B1	19911124		
F: AT, BE, CH, DE, FR, GR, LI, LU, NL, SE				
AT 97741	E	19931119	AT 1989-101406	19890319
US 5140048	A	19930618	US 1989-008654	19901030
US 5126442	A	19930630	US 1991-008735	19910108
US 5054593	A	19931019	US 1991-008739	19911110
JF 08172813	A2	19930713	JP 1991-116577	19911110
US 5356895	A	19941018	US 1991-008741	19910607
WO 9313775	A1	19930722	WO 1993-008746	19930115
W: AU, CA, JP				
FW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
AU 9335840	A1	19930803	AU 1993-008740	19930115
US 5811075	A	19980922	US 1995-487398	19950607
WO 9640663	A1	19961219	WO 1996-008776	19960607
W: AU, CA, IL, JP				
FW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
AU 9061586	A1	19961110	AU 1996-01586	19960607
US 5851099	A	19981212	US 1997-774861	19980110
US 6114323	A	20000905	US 1998-015612	19981217
US 2001115724	A1	20020822	US 2001-0084514	20010317
PRIORITY APPLN. INFO.:				
			US 1984-590810	A2 19840319
			US 1985-708032	A1 19851114
			US 1987-110958	A2 19871113
			US 1988-164930	A2 19881102
			US 1990-605654	A2 19901030
			US 1992-889141	A3 19920527
			US 1994-174343	B2 19940713
			EP 1989-101406	A 19890319
			US 1990-007747	B2 19900912
			US 1987-01534	A3 19870903
			US 1986-128504	B2 19860713
			US 1989-453835	A3 19891030
			US 1989-453838	E1 19891030
			US 1990-481869	A2 19901030
			US 1990-008415	A3 19901031
			US 1991-704487	E1 19910603
			US 1992-801318	A 19920117
			US 1992-878437	E1 19920525
			WO 1993-US750	A 19930115
			US 1993-161840	E1 19931003
			US 1994-150660	B1 19940915
			US 1995-487159	A 19950607

WO 1996-US9376 W 19960607  
 US 1997-784861 A1 1997 0116  
 US 1998-215612 A1 19981217  
 US 2000-561541 A3 20000423

OTHER SOURCE(S): MARPAT 126:122831

AB The present invention relates to compos., compns. and methods for inhibiting nonenzymic **crosslinking** (protein aging). Accordingly, a compn. is disclosed which comprises a di- or tri-aminoquinidine capable of inhibiting the formation of advanced glycosylation end products of target proteins. The method comprises contacting the target protein with the compn. Both industrial and therapeutic applications for the invention are envisioned, as food spoilage and animal protein aging can be treated.

IT **13431-34-0**, 4-Ethyl-3-thiosemicarbazide  
 RL: RCT (Reactant); FACT (Feactant or reagent)  
 (di- and triaminoquinidines and methods of use to prevent protein aging)

RN 13431-34-0 HCAPLUS  
 CN Hydrazinecarbothioamide, N-ethyl- (PCI) (CA INDEX NAME)

ETHN C NH NH2

L9 ANSWER 11 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1997:127504 HCAPLUS

DOCUMENT NUMBER: 126:129000

TITLE: Semicarbazide-containing linker compounds for formation of stably-linked **conjugates** and methods related thereto

INVENTOR(S): Berninger, Ronald W.; Lodge, Mark S.; Tarnowski, Stanley Joseph, Jr.

PATENT ASSIGNEE(S): Cellpro, Incorporated, USA

SOURCE: ECT Int. Appl., 33 pp.

CODEN: PIMED2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY APP. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9640163	A2	19961219	WO 1996-US8983	19960604
WO 9640163	A3	19970417		

W: JP  
 RW: AT, BE, CH, DE, DK, ES, FI, FR, GE, GR, IE, IT, LU, MC, NL, PT, SE  
 US 1356571 A 19990105 US 1995-486980 19950607  
 US 1995-486989 19950607

PRIORITY APPL. INFO.:

OTHER SOURCE(S): MARPAT 126:129000

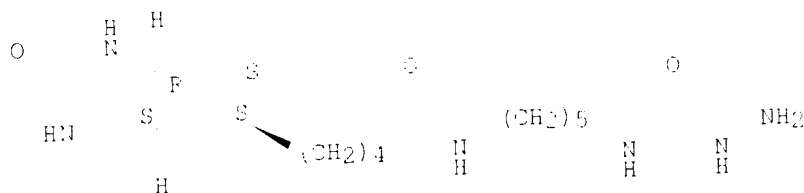
AB Linker compds. for formation of stably-linked **conjugates** are disclosed. Such linker compds. are semicarbazide-contg. linker compds. useful in forming **conjugates** having stable semicarbazone linkages. The stably-linked **conjugates** have utility in a variety of immunodiagnostic and sepn. techniques.

IT **186422-63-9P**  
 FL: PNU (Preparation, unclassified); RCT (Reactant); PREP (Preparation);  
 FACT (Feactant or reagent)

(semicarbazide-contg. linker compds. for formation of stably-linked  
**conjugates** and methods related thereto)

RN 186422-63-9 HCAPLUS  
 CN 1H-Thieno[3,4-d]imidazole-4-pentanamide, N-[5-  
 [(hydrazinocarbonylamino)pentyl]hexahydro-2-oxo-, [3aS-  
 (3a.alpha.,4.beta.,5a.alpha.)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L9 ANSWER 12 OF 23 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1997:2948 HCAPLUS

DOCUMENT NUMBER: 126:128771

TITLE: Substituted thioureas as **bifunctional**  
 chelators, their preparation, **conjugates**  
 with peptides, proteins, and antibodies, and their use  
 in imaging of tumors and thrombi

INVENTOR(S): Coughlin, Daniel J.; Belinka, Jr Benjamin A.

PATENT ASSIGNEE(S): Cytogen Corporation, USA

SOURCE: U.S., 32 pp., Cont.-in-part of U.S. 5,326,856.

CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5585468	A	19961217	US 1994-204197	19940627
US 5326856	A	19940705	US 1992-866375	19920409
WO 9301151	A1	19931028	WO 1993-US3208	19930408

W: CA, JP, US

FW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE

PRIORITY APPLN. INFO.:

US 1992-866375 19920409  
 WO 1993-US3208 19930408

OTHER SOURCE(S): MAEPAT 126:128771

GI

HQ2C

NHCSNHNHCOCH<sub>2</sub>NMe<sub>3</sub><sup>+</sup>Cl<sup>-</sup>

NHCSNHNHCOCH<sub>2</sub>NMe<sub>3</sub><sup>+</sup>Cl<sup>-</sup>

I

AB Chelating agents useful for coupling metal ions to biol. active mols. are  
 disclosed. In particular, substantial thioureas for chelating metals,  
 e.g. technetium, are provided that can be **conjugated** to a

targeting mol. such as an antibody, a peptide or a protein. Prepn. of the chelating agents of the invention, e.g. 1, is described, as are conjugation to an antibody and to a peptide and use of the conjugates in tumor imaging and thrombus imaging.

IT 6610-29-3, 4-Methyl-3-thiosemicarbazide

RL: PCT (Reactant); PACT (Reactant or reagent)

(reaction; substituted thioureas as **bifunctional** chelators, prepn., **conjugates** with peptides, proteins, and antibodies, and use in imaging of tumors and thrombi)

RN 6610-29-3 HCAPLUS

CN Hydrazinecarbothioamide, N-methyl- (9CI) (CA INDEX NAME)

S

MeNH C NH NH<sub>2</sub>

LS ANSWER 13 OF 38 HCAPLUS COPYRIGHT 1993 ACS

ACCESSION NUMBER: 1996:724187 HCAPLUS

DOCUMENT NUMBER: 116:4221

TITLE: Method of photochemical **immobilization** of ligands using quinones

INVENTOR(S): Jacobsen, Mogens Havsteen; Koch, Troels

PATENT ASSIGNEE(S): Jacobsen, Mogens, Havsteen, Den.

SOURCE: PCT Int. Appl., 98 pp.

CODEN: PIMXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9651597	A1	19961010	WO 1996-DK167	19960403
W:	AL, AM, AT, AU, AC, BE, BG, BR, BY, CA, CH, CN, CT, DE, DK, EE, ES, FI, GE, GR, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LF, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, PG, PU, SD, SE, SG, SI			
FW:	FE, IS, MW, SD, SE, SG, AT, BE, CH, DE, DK, ES, FI, FE, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN			
CA 2219082	AA	19961010	CA 1996-2217053	19960403
AU 9653329	A1	19961023	AU 1996-53329	19960403
AJ 698321	B2	19981203		
EP 820483	A1	19960128	EP 1996-909990	19960403
EP 820483	B1	20001213		
F:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI			
JP 11505554	T2	19990511	JP 1996-509895	19960403
JP 3124037	B2	20010115		
AT 198079	E	20001215	AT 1996-909990	19960403
ES 2153097	T3	20010216	ES 1996-909990	19960403
US 6933784	A	20000307	US 1997-08623	19971007
PRIORITY APPL. INFO.:			DK 1995-425	A 19960407
			WO 1996-DK167	W 19960403

OTHER SOURCE(S): CASREACT 116:4221; MARPAT 116:4221

AB A method is disclosed for **immobilizing** a ligand on the surface of a carbon-contg. substrate material, said method comprising a photochem. step of linking .gtoreq.1 photochem. reactive compds. to a carbon-contg.



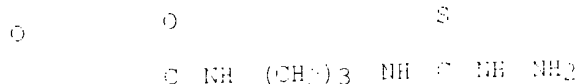
material surface, wherein the photochem. reactive compd. is a quinone compd. contg. a cyclic hydrocarbon or 2-10 fused cyclic hydrocarbons, with at least 2 **conjugated** carbonyl groups, and wherein the photochem. step comprises irradiation of the photochem. reactive compd. with nonionizing electromagnetic radiation having a wavelength in the range from UV to visible light. The products of this invention can be used as, e.g., carriers for solid-phase immunoassays.

IT 172422-03-6P

EL: EIT (Reactant); SYN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(photochem. **immobilization** of ligands using quinones)

RN 172422-03-6 HCAPLUS

CN 2-Anthracenecarboxamide, N-[3-[(hydrazinothioxomethyl)amino]propyl]-9,10-dihydro-9,10-dioxo- (PCI) (CA INDEX NAME)



O

L9 ANSWER 14 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1995:319701 HCAPLUS  
DOCUMENT NUMBER: 123:89553  
TITLE: PEG hydrazone and PEG oxime linkage forming reagents and protein derivatives.  
INVENTOR(S): Wright, David E.  
PATENT ASSIGNEE(S): Ortho Pharmaceutical Corp., USA  
SOURCE: Eur. Pat. Appl., 47 pp.  
CODEN: EPXMLW  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 605963	A2	19940713	EP 1993-309825	19931709
EP 605963	A3	19951103		
FI: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
CA 2110543	AA	19940610	CA 1993-2110543	19931701
FI 9404495	A	19940610	FI 1993-5485	19931208
NO 9304477	A	19940610	NO 1993-4477	19931208
ZA 9309314	A	19950608	ZA 1993-9214	19931208
AU 9252392	A1	19940623	AU 1993-52385	19931209
JP 7196925	A2	19950801	JP 1993-340709	19931209
PRIORITY APPL. INFO.:			US 1992-987739	19921209
			US 1993-45051	19930407
			US 1993-157343	19931123

AB Comp. is. for modifying polypeptides with PEG or other water-sol. org. polymers are described. The water-sol. polymer reagents include hydrazine, hydrazine carboxylate, semicarbazide, thiosemicarbazide, carbonic acid dihydrazide, carbazide, thiocarbazide, and arylhydrazide derivs. as well as oxylamine derivs. of water-sol. org. polymers, such as

polyethylene glycol, polypropylene glycol, polyoxyethylated polyol, heparin, heparin fragments, dextran polysaccharides, polyamino acids, and polyvinyl alc. Kits for modifying polypeptides with the above water-sol. polymer reagents are also provided. Thus, erythropoietin was modified by oxim. and treatment with monomethoxypolyoxyethylene semicarbazide and the product was sepd. by chromatog. The antigenicity and the effect on hematocrit levels of the above derivs. were demonstrated.

IT 160556-27-4DP, reaction products with protein derivs.  
RL: BAC (Biological activity or effector, except adverse); RSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)  
(pregn. and biol. activity of polyoxyethylene-coupled protein derivs.)

RN 160556-27-4 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-[2-[(hydrazinocarbonyl)amino]ethyl]-.omega.-hydroxy- (9CI) (CA INDEX NAME)

Q

HO  $\text{CH}_2 \text{CH}_2 \text{O} \text{CH}_2 \text{CH}_2 \text{NH} \text{C} \text{NH} \text{NH}_2$   
n

IT 160556-27-4P 160556-28-5P  
RL: RCT (Reactant); SEN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(pregn. and biol. activity of polyoxyethylene-coupled protein derivs.)

RN 160556-27-4 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-[2-[(hydrazinocarbonyl)amino]ethyl]-.omega.-hydroxy- (9CI) (CA INDEX NAME)

Q

HO  $\text{CH}_2 \text{CH}_2 \text{O} \text{CH}_2 \text{CH}_2 \text{NH} \text{C} \text{NH} \text{NH}_2$   
n

RN 160556-28-5 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-[2-[(hydrazinothioxomethyl)amino]ethyl]-.omega.-hydroxy- (9CI) (CA INDEX NAME)

S

HO  $\text{CH}_2 \text{CH}_2 \text{O} \text{CH}_2 \text{CH}_2 \text{NH} \text{C} \text{NH} \text{NH}_2$   
n

L9 ANSWER 15 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1994:95783 HCAPLUS

DOCUMENT NUMBER: 120:95783

TITLE: Inhibitors of thrombosis

INVENTOR(S): Vlasuk, Georg Phillip; Webb, Thomas Roy; Pearson, Daniel Andrew

PATENT ASSIGNEE(S): Corvas International, Inc., USA

SOURCE: PCT Int. Appl., 37 pp.

CODEN: PIRX82

DOCUMENT TYPE: Patent

LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 4  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9313756	A1	19930819	WO 1-93-US1307	19930812
W: CA, JP				
FW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 617039	A1	19941114	EP 1-93-905930	19930112
EP 617019	B1	19980930		
F: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
JF 07583961	T2	19950427	JF 1993-514315	19930112
JF 3194953	B2	20010806		
AT 171709	E	19981015	AT 1993-305930	19930212
CA 2124339	C	20020910	CA 1993-2129339	19930212
PRIORITY APPLN. INFO.:			US 1992-836123	A 19920214
			WO 1993-US1307	W 19930212

OTHER SOURCE(S): MAPPAT 120:95783

AB Peptide aldehyde analogs, AcR-AA-L-Pro-Arg-al (AcR = hydrophobic acyl group; AA = Glu, Asp, or equiv.), inhibit thrombin or Factor Xa and are thus useful for preventing or treating conditions in mammals characterized by abnormal thrombosis. N-(2-phenylpropyl)-L-Asp-L-Pro-L-argininal (prepn. given) inhibited thrombin, Factor Xa, and plasmin with IC50 values of 234, 91.5, and 326 nM, resp., and showed antithrombotic activity in a rat model.

IT 139976-29-7P 151275-26-2P

RL: FCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); FACT (Reactant or reagent)  
(prepn. and reaction of, in prepn. of antithrombotic peptide aldehyde analog)

RN 139976-29-7 HCAPLUS

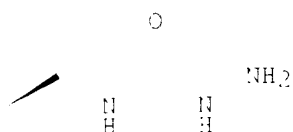
CN Cyclohexanecarboxylic acid, 4-[[[(hydrazinocarbonyl)amino]methyl]-, trans-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 139976-28-6

CME C9 H17 N3 O3

Relative stereochemistry.



HO2C

CM 2

CRN 76-05-1

CME C2 H F3 O2

F

F C CO<sub>2</sub>H

F

RN 111275-25-2 HCAPLUS  
 CN Hydrizinecarboxamide, N-(diphenylmethyl)-, mono(trifluoroacetate) (9CI)  
 (CA INDEX NAME)

CM 1

CPN 110908-39-7  
 CMF C14 H15 N3 O

O

H<sub>2</sub>N NH C NH CHPh<sub>2</sub>

CM 2

CPN 76-05-1  
 CMF C2 H F3 O2

F

F C CO<sub>2</sub>H

F

L9 ANSWER 16 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1994:49591 HCAPLUS  
 DOCUMENT NUMBER: 120:49591  
 TITLE: Substituted thioureas as **bifunctional**  
 chelators for **conjugation** to antibodies or  
 other biological targeting molecules  
 INVENTOR(S): Coughlin, Daniel J.; Belinka, Benjamin A., Jr.  
 PATENT ASSIGNEE(S): Cytogen Corp., USA  
 SOURCE: PCT Int. Appl., 89 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 93/1151	A1	19931018	WO 1993-US3208	19930408
W: CA, JP, US				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
US 5326356	A	19940705	US 1992-366375	19920409
EP 635001	A1	19950125	EP 1993-911594	19930408

EP 635001 B1 19970823  
 E: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE  
 JP 68102340 T2 19960311 JP 1993-518423 19930403  
 AT 117044 E 19970911 AT 1997-911594 19980403  
 ES 311495 T3 19950116 ES 1995-911594 19960403  
 US 5544668 A 19961217 US 1994-204197 19940623  
 US 5519835 A 19960515 US 1994-268443 19940630  
 PRIORITY APPLN. INFO.: US 1993-06375 19930403  
 WO 1993-003208 19930403

OTHER SOURCE(S): MARPAT 120:49591

AB The title chelating agents are LD[NHC(S)NHR]<sub>2</sub> (L = linker; D = (cyclic alkyl, aryl; R = H, (NH)a(CH<sub>2</sub>)<sub>b</sub>(C(Y))<sub>c</sub>NH-d(CH<sub>2</sub>)<sub>e</sub>Z (a = 0, 1; b, e = 0-10; c = 0, 1 (if c = 1, Y = S, O, H<sub>2</sub>); d = 0-2; Z = H, SO<sub>3</sub>H, CO<sub>2</sub>H, OH, H<sub>2</sub>PO<sub>3</sub>, R'(R')<sub>3</sub>X- (R' = C1-4 alkyl; X- = counterion, such as halide or acid anion)); the chelating agents are useful for coupling metal ions to biol. active mol's. (antibodies, peptides, etc.). Prepn. of several chelating agents of the invention is described. Thus, 3,5-di-(1-trimethylammoniumacetyl)-4-thiosemicarbazidebenzoic acid dichloride salt (I) was prepd. from 3,5-diisothiocyanatobenzoic acid (prepn. given) and (carboxymethyl)trimethylammonium chloride hydrate. I was **conjugated** to a peptide (SYEGDLVEGDF-NH<sub>2</sub>), and the **conjugate** was labeled with <sup>99m</sup>Tc. The labeled peptide **conjugate** was used in the imaging of thrombi in rabbits. Prepn. and use in tumor imaging of a labeled antibody **conjugate** is also described.

IT 6610-29-3, 4-Methyl-3-thiosemicarbazide  
 RL: PCT (Peactant); RACT (Reactant or reagent)  
 (reaction of, in **bifunctional** substituted thiourea chelating agent prepn.)

RN 6610-29-3 HCAPLUS

CN Hydrazinecarbothioamide, N-methyl- (9CI) (CA INDEX NAME)

3

MeNH C NH NH<sub>2</sub>

L9 ANSWER 17 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1993:617414 HCAPLUS  
 DOCUMENT NUMBER: 119:117414  
 TITLE: Peptide aldehyde analogs for trypsin inhibitors  
 INVENTOR(S): Brunck, Terence Kevin; Pepe, Michael Gary; Pearson, Daniel Andrew; Webb, Thomas Foy  
 PATENT ASSIGNEE(S): Corvas International, Inc., USA  
 SOURCE: PCT Int. Appl., 61 pp.  
 CODEN: PEXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9314779	A1	19930605	WO 1993-US906	19930129
W: CA, JP				
FW: AT, BE, CH, DE, DK, ES, FF, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 627925	A1	19941214	EP 1993-905778	19930129
E: AT, BE, CH, DE, DK, ES, FF, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				

JP 07503715	T2	19950420	JP 1993-513488	19930129
US 5534498	A	19960709	US 1993-11666	19930129
PRIORITY APPLN. INFO.:			US 1992-828388	19920130
			US 1993-11666	19930129
			WO 1993-US906	19930129

OTHER SOURCE(S): MARPAT 119:217414

AB Peptide aldehyde analogs are disclosed which have substantial potency and specificity as inhibitors of mammalian pancreatic trypsin. The compds. of the invention are useful in the prevention and treatment of tissue damage or destruction assocd. with pancreatitis. Prepn. of the analogs is described. Thus, N-t-butoxycarbonyl-L-Asp-L-Pro-L-argininal (I) (prepn. given; had a  $K_i$  against trypsin of 0.00045  $\mu$ M. The effectiveness of I in an animal model for pancreatitis was also demonstrated.

IT **139976-29-7P 150908-39-7P**

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(prepn. and reaction of, in peptide aldehyde analog prepn. for trypsin inhibitor)

RN 139976-29-7 HCAPLUS

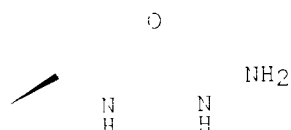
CN Cyclohexanecarboxylic acid, 4-[[[(hydrazinocarbonyl)amino]methyl]-, trans-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CEN 139976-28-6

CMF C9 H17 N3 O3

Relative stereochemistry.



HO2C

CM 2

CEN 76-05-1

CMF C2 H F3 O1

F

F C CO2H

F

RN 150908-39-7 HCAPLUS

CN Hydrazinecarboxamide, N-(diphenylmethyl)- (9CI) (CA INDEX NAME)

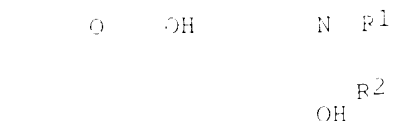
G

H2N NH C NH CHPh2

L9 ANSWER 19 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1992:214841 HCAPLUS  
 DOCUMENT NUMBER: 116:214841  
 TITLE: Preparation of anthracycline immunoconjugates  
 as neoplasm inhibitors  
 INVENTOR(S): Kaneko, Takushi; Willner, David; Monkovic, Ivo;  
 Greenfield, Robert S.; Braslawsky, Gary R.  
 PATENT ASSIGNER(S): Bristol-Myers Squibb Co., USA  
 SOURCE: Eur. Pat. Appl., 45 pp.  
 CODEN: EPMXDW  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 457250	A1	19911121	EP 1991-107737	19910513
EP 457250	A3	19910701		
EP 457250	B1	19910714		
F: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE				
US 5137377	A	19930811	US 1990-522996	19900514
US 5137377	B1	19930130		
AU 2174538	A1	19911114	AU 1991-74038	19910403
AU 648850	B2	19940310		
FI 9102285	A	19911115	FI 1991-2285	19910510
JP 04350765	A2	19911207	JP 1991-199757	19910510
JP 3010319	B2	20000211		
ZA 9103591	A	19920126	ZA 1991-3591	19910513
AT 182141	E	19930715	AT 1991-107737	19910513
ES 1134761	T3	19911016	ES 1991-107737	19910513
CA 1941503	AA	19911115	CA 1991-2042503	19910514
CA 1941503	C	20020713		
US 5349066	A	19940410	US 1992-885062	19920408
JP 1000026404	A2	20000115	JP 1999-131583	19990512
JP 3234980	B2	20011104		

PRIORITY APPLN. INFO.: US 1990-522996 A 19900514  
 JP 1991-199757 A3 19910510  
 OTHER SOURCE(S): MARPAT 116:214841  
 GI



AB Anthracycline derivs. I [R<sup>1</sup> = NECONH(CH<sub>2</sub>)<sub>n</sub>SSR<sup>8</sup>, NHCONHNHCONH(CH<sub>2</sub>)<sub>n</sub>SSR<sup>8</sup>, NECSNH(CH<sub>2</sub>)<sub>m</sub>CH:CH(CH<sub>2</sub>)<sub>n</sub>SSR<sup>8</sup>, NHC<sub>2</sub>O<sub>2</sub>(CH<sub>2</sub>)<sub>n</sub>SSR<sup>8</sup>, NHArCONH(CH<sub>2</sub>)<sub>n</sub>SSR<sup>8</sup>, etc.; m, n = 1-10; R<sup>8</sup> = (substituted) 2-pyridyl, -phenyl; Ar = phenylene; R<sup>2</sup> = Me, CH<sub>2</sub>OH, CH<sub>2</sub>COO(CH<sub>2</sub>)<sub>3</sub>Me, CH<sub>2</sub>COCH(OEt)<sub>2</sub>; R<sup>3</sup> = OMe, OH, H; R<sup>4</sup> = NH<sub>2</sub>, NHCOCF<sub>3</sub>, 4-morpholinyl, 3-cyano-4-morpholinyl, 1-piperidinyl, NHC<sub>2</sub>H<sub>5</sub>Ph, N(CH<sub>2</sub>Ph)<sub>2</sub>, etc.; R<sup>5</sup> = OH, tetrahydropyranyloxy, H; R<sup>6</sup> = OH, H; R<sup>6</sup> noteq. OH when R<sup>5</sup> = OH or tetrahydropyranyloxy], related compds., and their **conjugates** with ligands and antibodies, were prepd. Thus, 1-amino-4-[(2-pyridinyl)dithio]-2-butene-HCl (prepn. given) was treated with di(2-pyridyl) thionocarbonate and the product formed was condensed with Me<sub>3</sub>CO<sub>2</sub>CNHNH<sub>2</sub>. Deprotection of the resulting product by CF<sub>3</sub>CO<sub>2</sub>H gave N-[4-(2-pyridinyl)dithio]-2-butenylhydrazinecarbothioamide. This was condensed with adriamycin-HCl to give adriamycin 13-N-4-[(2-pyridinyl)dithio]-2-butenylhydrazinecarbothioamide thiosemicarbazene.entdot.HCl (II). The **immunoconjugate** of II with thiolated monoclonal antibody 5E9 had IC<sub>50</sub> of 3.0 times. 101-7M against Burkitt's lymphoma cells.

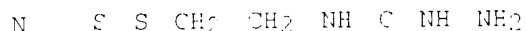
IT 133701-16-3P 140691-64-1P

RL: SPH (Synthetic preparation); PREP (Preparation)  
(prepn. of, as intermediate for anticancer **immunoconjugates**)

RN 133701-16-3 HCAPLUS

CN Hydrazinecarboxamide, N-[2-(2-pyridinyldithio)ethyl]- (9CI) (CA INDEX NAME)

O



RN 140691-64-1 HCAPLUS

CN Hydrazinecarbothioamide, N-[4-(2-pyridinyldithio)-2-butenyl]- (9CI) (CA INDEX NAME)



S

N S S CH<sub>2</sub> CH CH CH<sub>2</sub> NH C NH NH<sub>2</sub>

LE ANSWER 19 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1991:253927 HCAPLUS  
 DOCUMENT NUMBER: 114:253927  
 TITLE: New hydrazone derivatives of Adriamycin and their  
**immunoconjugates** - a correlation between acid  
 stability and cytotoxicity  
 AUTHOR(S): Kaneko, Takushi; Willner, David; Monkovic, Ivo; Knipe,  
 Jay O.; Braslawsky, Gary R.; Greenfield, Robert S.;  
 Vyas, Dolatrai M.  
 CORPORATE SOURCE: Bristol-Myers Squibb Co., Wallingford, CT, 06492-7660,  
 USA  
 SOURCE: Bioconjugate Chemistry (1991), 2(3), 133-41  
 CODEN: BOCHES; ISSN: 1043-1802  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB New N-substituted hydrazine linkers were synthesized and their hydrazone  
 derivs. of adriamycin were prep'd. The adriamycin derivs. were  
**conjugated** with a monoclonal antibody, 5E9. The release rate of  
 adriamycin from the hydrazones and from some of the **conjugates**  
 was studied, and their relationship to the cytotoxicity against 5E9-pos.  
 Daudi cells was investigated.  
 IT 133701-16-3P 133701-22-1P  
 EL: SPN (Synthetic preparation); PFEP (Preparation)  
 (prepn. and condensation of, with adriamycin, hydrazone from)  
 EN 133701-16-3 HCAPLUS  
 CN Hydrazinecarboxamide, N-[2-(2-pyridinyldithio)ethyl]- (9CI) (CA INDEX  
 NAME)

O

N S S CH<sub>2</sub> CH<sub>2</sub> NH C NH NH<sub>2</sub>

EN 133701-22-1 HCAPLUS  
 CN Hydrazinecarbothioamide, N-[4-(2-pyridinyldithio)-2-butenyl]-, (E)- (9CI)  
 (CA INDEX NAME)

Double bond geometry as shown.

N S S E H H  
 S N N NH<sub>2</sub>

S

L9 ANSWER 20 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1901:20183 HCAPLUS  
 DOCUMENT NUMBER: 114:20183  
 TITLE: Radiolabeling of protein with radioisotopes of copper  
 using p-carboxyalkylphenylglyoxal bis-(4N-methylthiosemicarbazone) (TSC) **bifunctional**  
 chelates  
 AUTHOR(S): McPherson, I. W.; Umbricht, G.; Knapp, E. F., Jr.  
 CONTRIBUTOR SOURCE: Health Saf. Res. Div., Oak Ridge Natl., Oak Ridge, TN,  
 37831-6012, USA  
 SOURCE: Journal of Labelled Compounds and Radiopharmaceuticals  
 (1990), 28(8), 877-89  
 CODEN: JLCR34; ISSN: 0362-4803  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 OTHER SOURCE(S): CASREACT 114:20183  
 GI

NNEC(S)NHMe

HO<sub>2</sub>C(CH<sub>2</sub>)<sub>n</sub>

C CF

NHMe(S)NHMe 1

AB A series of p-carboxyalkylphenylglyoxal and p-carboxyalkyl-1,2-diketobis-(N-methylthiosemicarbazone) **bifunctional** ligands I (R = H or Me, n = 1-9) were prepd. and evaluated for use in binding radioisotopes of Cu to antibodies. An improved synthesis of the requisite .alpha.-keto aldehyde and 1,2-diketone substrates used for derivatization to the bis-TSC **bifunctional** chelates was developed. This approach utilizes a modified Kornblum method and provides a simple alternative to the usual method for fabrication of the 1,2-bis ligands, which avoids the use of highly toxic SeO<sub>2</sub> for oxidn. of substituted acetophenones to 1,2 dicarbonyl compds. The overall yields of the bis-TSC chelates using this procedure were 8-60%. The effects of the alkyl chain length and substitution on the C-2 position on **bifunctional** chelates for attaching radioisotopes of copper to proteins were studied. Following complexing <sup>64</sup>Cu or <sup>67</sup>Cu to the bis chelate, the acid moiety of the chelate was activated as the tetrafluorophenyl ester. The copper-labeled activated chelate was attached to bovine serum albumin under mild conditions in 3% to 40% yield. The shorter chain analog of the chelates from the 1,2-diketones give the highest radiolabeling yields.

IT 6610-29-3

RL: RCT (Reactant); FACT (Reactant or reagent)  
 (reaction of, with carboxyalkylphenylglyoxal derivs.)

EN 6610-29-3 HCAPLUS

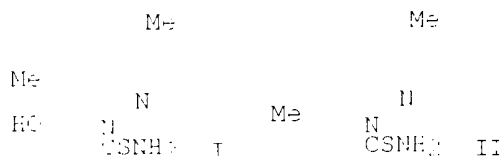
CU Hydrazinecarbothioamide, N-methyl- (9CI) (CA INDEX NAME)

?

MeNH C NH NH<sub>2</sub>

L9 ANSWER 21 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1988:152962 HCAPLUS

DOCUMENT NUMBER: 108:150351  
 TITLE: Reactions of 1,4-bifunctional derivatives of hydrazine with 1,2-diketones  
 AUTHOR(S): Zelenin, V. N.; Slobod, O. V.; Tomchin, A. B.  
 CORPORATE SOURCE: Uchen.-Med. Akad., Leningrad, USSR  
 SOURCE: Zhurnal Obshchei Khimii (1987), 57(3), 584-95  
 CODEN: ZOPHAA; ISSN: 0044-460X  
 DOCUMENT TYPE: Journal  
 LANGUAGE: Russian  
 OTHER SOURCE(S): CASREACT 1-8:150352  
 GI



AB Hydrazine derivs., e.g., aminoguanidine nitrate, PhNHCONHNH<sub>2</sub>, amidrazonium  
 azides, R<sub>1</sub>HNCSNHNH<sub>2</sub> (R<sub>1</sub> = H, Me, Et) condense with RCOCH<sub>2</sub>CO<sub>2</sub>R (R = Me, Ph)  
 to give, depending on reaction conditions, 5-hydroxy- and  
 5-hydrazino-3-pyrazolines, mono- and bis(hydrazones), and also the  
 corresponding pyrazoles. Thus, treating MeCOCH<sub>2</sub>COMe with H<sub>2</sub>NCSNHNH<sub>2</sub> gave  
 pyrazoline I which dehydrated in refluxing solvent to give the  
 corresponding pyrazole II. Anal. obtained was R<sub>1</sub>NHN:CRCH<sub>2</sub>CR:NNHR<sub>1</sub> [R =  
 Me, R<sub>1</sub> = CONHPh, C(=NH)(NH<sub>2</sub>).H<sub>2</sub>O<sub>3</sub>].

IT 6610-29-3, 4-Methyl-3-thiosemicarbazide 13431-34-0,  
 4-Ethyl-3-thiosemicarbazide  
 FL: PCT (Reactant); EACT (Reactant or reagent)  
 (condensation and cyclodehydration of, with diketones)  
 RN 6610-29-3 HCAPLUS  
 CN Hydrazinecarbothioamide, N-methyl- (901) (CA INDEX NAME)

S

MeNH C NH NH<sub>2</sub>

RN 13431-34-0 HCAPLUS  
 CN Hydrazinecarbothioamide, N-ethyl- (901) (CA INDEX NAME)

S

EtNH C NH NH<sub>2</sub>

L9 ANSWER 22 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1987:196867 HCAPLUS  
 DOCUMENT NUMBER: 106:196867  
 TITLE: Polymers containing the [2H]-1,2,4-triazoline-3-thione  
 ring  
 AUTHOR(S): Patrisky, Alan E.; Cato, Stephen J.; Heilmann, Steven  
 M.; Rasmussen, Jerald K.; Krepski, Larry R.

CORPORATE SOURCE: Chem. Dep., Univ. Florida, Gainesville, FL, 32611, USA  
 SOURCE: Journal of Polymer Science, Part A: Polymer Chemistry  
 (1987), 25(1), 311-26  
 CODEN: JPACEC; ISSN: 0887-624X  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB High-mol.-wt. polymers contg. [2H]-1,2,4-triazoline-3-thione rings are  
 prep'd. by the condensations of diisothiocyanates with bis(acid hydrazides)  
 to give intermediate polymeric acylthiosemicarbazides that are ring-closed  
 by refluxing in 1M aq. sodium carbonate. Thermal cyclization of the  
 polymeric acylthiosemicarbazides leads to **crosslinked** insol.  
 products. The acylation of bis(thiosemicarbazides) with bis(acid  
 chlorides) produces polymers of a similar structure but lower mol. wt.  
 IT **6610-31-7P**, 4-Butylthiosemicarbazide **13431-41-9P**,  
 4-Benzylthiosemicarbazide  
 FL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
 (Reactant or reagent)  
 (prepn. and reaction with Et imide ester hydrochlorides)  
 EN 6610-31-7 HCAPLUS  
 CN Hydrazinecarbothioamide, N-butyl- (9CI) (CA INDEX NAME)

S

CCCCNC(=O)NNH2

EN 13431-41-9 HCAPLUS  
 CN Hydrazinecarbothioamide, N-(phenylmethyl)- (9CI) (CA INDEX NAME)

S

NC(=O)NC(=O)C1=CC=CC=C1

IT **108144-98-5P**  
 FL: SPN (Synthetic preparation); PREP (Preparation)  
 (prepn. of)  
 EN 108144-98-5 HCAPLUS  
 CN 1,4-Benzenedicarbonyl dichloride, polymer with N,N'-1,6-  
 hexanediylbis[hydrazinecarbothioamide] (9CI) (CA INDEX NAME)

CM 1

CRN 56473-15-5  
 CMF C8 H20 N6 S2

S

S

NC(=O)NC(=O)C1CCC(CC1)NC(=O)NNH2

CM 2

CRN 100-20-9  
 CMF C8 H4 Cl2 O2

C  
C C1

C1 C

O

IT 56473-15-5, 1,6-Hexanabis(thiosemicarbazide)  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (reaction of, with Et benzimidazole hydrochloride)  
 RN 56473-15-5 HCAPLUS  
 CH Hydrazinecarbothioamide, N,N'-1,6-hexanediylbis- (9CI) (CA INDEX NAME)

S S

H<sub>2</sub>N NH C NH (CH<sub>2</sub>)<sub>6</sub> NH C NH NH<sub>2</sub>

L# ANSWER 23 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1986:520711 HCAPLUS  
 DOCUMENT NUMBER: 105:120711  
 TITLE: Search for technetium-99m labeled DTS  
**bifunctional** radiopharmaceutical: role of  
 functional groups in myocardial accumulation  
 AUTHOR(S): Hosotani, Takeo; Yokoyama, Akira; Arano, Yasushi;  
 Horiuchi, Kazuko; Saji, Hideo; Torizuka, Kanji  
 CORPORATE SOURCE: Fac. Pharm. Sci., Kyoto Univ., Kyoto, 606, Japan  
 SOURCE: Applied Radiation and Isotopes (1986), 37(6), 505-11  
 CODEN: ARISEF; ISSN: 0383-2889  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 OTHER SOURCE(S): CASREACT 105:120711

AB Various mois. contg. a neutral 99mTc-dithiosemicarbazone (DTS) structure  
 as the Tc chelating site, along with various functional groups (NH<sub>2</sub>, CO<sub>2</sub>H  
 or iso-Bu group with diverse charge) were tested for their chem. or biol.  
 functions. The study on the effect of those functional groups was carried  
 out in vitro and in vivo. The validity of introducing an NH<sub>2</sub> group along  
 with the Tc chelating site DTS for myocardial accumulation is discussed.

IT 6610-29-3  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (reaction of, with phenylglyoxals)  
 RN 6610-29-3 HCAPLUS  
 CH Hydrazinecarbothioamide, N-methyl- (9CI) (CA INDEX NAME)

C

MeNH C NH NH<sub>2</sub>

L# ANSWER 24 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1986:438342 HCAPLUS

DOCUMENT NUMBER: 105:38342  
 TITLE: Synthesis and evaluation of a new **bifunctional** chelating agent for technetium-99m labeling proteins: p-carboxyethylphenylglyoxal-di(N-methylthiosemicarbazone)  
 AUTHOR(S): Arano, Yasushi; Yokoyama, Akira; Magata, Yasuhiro; Saji, Hideo; Horikoshi, Kazuko; Torizuka, Kanji  
 CORPORATE SOURCE: Fac. Pharm. Sci., Kyoto Univ., Kyoto, 606, Japan  
 SOURCE: International Journal of Nuclear Medicine and Biology (1986), 12(6), 429-30  
 CODEN: IJNMCI; ISSN: 0947-0749  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 GI

HQ2CCH2CH2 C NNBOSNHMe  
 HC NNBOSNHMe I

AB A new **bifunctional** chelating agent, p-carboxyethylphenylglyoxal-di(N-methylthiosemicarbazone) (I), contg. a di(N-methylthiosemicarbazone) as the Tc coordinating site and an aralkyl carboxylate site for the protein **conjugation** was synthesized. Coupling to human serum albumin (HSA), selected as a model protein, was carried out by the phosphorylazide method using diphenylphosphoryl azide (DPPA). The **conjugation** level of I to HSA played a crit. role in its biol. evaluation. A 99mTc-I-HSA with high in vivo stability was obtained when I was coupled to HSA at 1:1 molar ratio. This compd. showed similar in vivo stability to 131I-labeled HSA in mice and rabbits.

IT 6610-29-3  
 RL: PRP (Properties)  
 (conjugation of, with acetylphenylpropionic acid)  
 EN 6610-29-3 HCAPLUS  
 CN Hydrazinecarbothioamide, N-methyl- (9CI) (CA INDEX NAME)

3

MeNH C NH NH2

L9 ANSWER 25 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1985:184835 HCAPLUS  
 DOCUMENT NUMBER: 102:184835  
 TITLE: p-Glyoxalphenylalkylcarboxylic acid bis(thiosemicarbazone) derivatives  
 PATENT ASSIGNEE(S): Nihon Med.-Physics Co., Ltd., Japan  
 SOURCE: Jpn. Kokai Tokkyo Koho, 6 pp.  
 CODEN: JPKXAF  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY ACC. NUM. COUNT: .  
 PATENT INFORMATION:

PATENT NO. KINL DATE APPLICATION NO. DATE

JP 59193870	A2	19841102	JP 1983-68850	19830419
JP 04016465	B4	19900324		
AU 6319934	A1	19841005	AU 1983-19934	19831006
AU 661558	B2	19870514		
US 4558211	A	19851217	US 1983-533884	19831037
CA 1200476	A1	19860701	CA 1983-438615	19831037

PRIORITY APPLN. INFO.:

JP 1983-68340 19830419  
JP 1983-68351 19830419

OTHER SOURCE(S): CASREACT 1:2:184835

AB **Bifunctional** ligand title derivs. 4-HO<sub>2</sub>C(CH<sub>2</sub>)nC<sub>6</sub>H<sub>4</sub>C(:NNHCSNHMe)CH:NNHCSNHMe I (n = 1-4) were prepd. by reaction of 4-HO<sub>2</sub>C(CH<sub>2</sub>)nC<sub>6</sub>H<sub>4</sub>COCHO (II) with H<sub>2</sub>NNHCSNHMe (III). I are useful as radioactive diagnostic reagents labeled with radioactive metals. Thus, refluxing 1.76 g 4-HO<sub>2</sub>CCH<sub>2</sub>OC<sub>6</sub>H<sub>4</sub>COMe with 1.22 g SeO<sub>2</sub> in dioxane 7 h gave II (n = 1), which (in EtOH) was added to 2.1 g III in 15 mL N aq. HCl at 60-degrees. to ppt. 1.1 g I (n = 1).

IT **6610-29-3**

RL: PCT (Reactant); FACT (Reactant or reagent)  
reaction of, with phenylglyoxal derivs.:

RN 6610-29-3 HCAPLUS

CN Hydrazinercarbothioamide, N-methyl- 9CI) (CA INDEX NAME)

S

MeNH C NH NH<sub>2</sub>

L9 ANSWER 26 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1985:25570 HCAPLUS

DOCUMENT NUMBER: 102:25570

TITLE: Basic polymers

PATENT ASSIGNEE(S): Asahi Chemical Industry Co., Ltd., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 6 pp.

CODEN: JKXKAF

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 59152905	A2	19840831	JP 1983-26392	19830222
JP 04069169	B4	19921105		

PRIORITY APPLN. INFO.:

JP 1983-26892 19830222

GI

CHCH<sub>2</sub>

CHCH<sub>2</sub>

R<sup>3</sup>

E1

N

CHCH<sub>2</sub>

I

R<sup>2</sup>

II

N N

III

AB The 2-50:50-73 (molar) I-II copolymers substituted with 10-36 mol% (based on total benzene rings) nuclear -COX group (X = OH, Cl) were treated with (methyl)thiosemicarbazide, an alkali, and then a nitrite salt in HNO<sub>3</sub> to obtain the **crosslinked** title polymers having 10-32 mol% (based on total benzene rings) III groups (on benzene rings), useful for anion exchangers (R's = H, Cl-4 hydrocarbyl). Thus, 17:33 m-divinylbenzene-styrene copolymer was subjected to a Friedel-Crafts reaction with oxalyl chloride in CCl<sub>2</sub> to obtain a chlorocarbonyl deriv. (I, 53 mol% COCl), which (11.54 g) was mixed with 150 mL EtOH and 21.0 g methylthiosemicarbazide, stirred under reflux for 2 h, filtered, washed with acetone-H<sub>2</sub>O, heated with 60 g NaOH in 300 mL water at 100.degree. for 1.5 h, filtered, washed with water, suspended in 100 mL water, and treated with 0.2 g NaNO<sub>2</sub> and 50 mL concd. HNO<sub>3</sub> at 45.degree. for 2 h to give 13.345 g polymer (52 mol % 4-methyltriazole group) having exchange capacity (HCl form) 1.71 mequiv/g.

IT 6610-29-3

RI: USES (Uses)

(on triazole group-contg. styrene deriv. polymer anion exchanger  
manuf.)

RI 6610-29-3 HCAPLUS

CH Hydrazinecarbothioamide, N-methyl- (9CI) (CA INDEX NAME)

S

MeNH C NH NH<sub>2</sub>

19 ANSWER 27 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1981:463041 HCAPLUS

DOCUMENT NUMBER: 95:68081

TITLE: 1-Oxopropionaldehydebis(thiosemicarbazone) derivatives

INVENTOR(S): Yakoyama, Akira; Arano, Yasushi

PATENT ASSIGNOR(S): Nihon Medel-Physics Co., Ltd., Japan

SOURCE: Eur. Pat. Appl., 20 pp.

CODEN: EPXXOW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 24464	A1	19810311	EP 1980-190199	19800110
EP 24464	B1	19810512		
E: BE, DE, FR, GB, NL, SE				
JP 56034664	A1	19810406	JP 1979-110821	19790829
JP 56034684	A1	19810406	JP 1979-110822	19790829
AU 8054731	A1	19810305	AU 1980-54721	19800118
AU 517412	B1	19810305		
US 42-7362	A	19810301	US 1980-113341	19800118
CA 1175418	A1	19810301	CA 1980-347997	19800118
US 4338246	A	19810706	US 1980-117947	19800314
PRIORITY APPLN. INFO.:			JP 1979-110821	19790829
			JP 1979-110822	19790829
			US 1980-113341	19800118

AB A radiolabeled diagnostic agent prepd. from a protein and a radioactive element and a **bifunctional** chelating agent is quite stable. The



chelating agent 3-carboxy-2-oxopropionaldehyde bis(N-methylthiosemicarbazone) (I) [78277-80-2], prep'd. from Et diethoxyacetate [10425-09-7] and N-methylthiosemicarbazide [6610-29-3] and hydrolysis of the resulting 3-ethoxycarbonyl-2-oxopropionaldehyde bis(N-methylthiosemicarbazone) [78277-81-5], was converted to a mixed anhydride by treatment with iso-Bu chloroformate. Human serum albumin was mixed with the anhydride and subjected to dialysis followed by lyophilization. The albumin-I complex was treated with  $^{99m}\text{Tc}$  (10.5mCi) at pH 8.5 in the form of a pertechnetate and reduced with  $\text{SnCl}_2$  soln. to yield a  $^{99m}\text{Tc}$ -albumin-I complex useful as a radioactive diagnostic agent. The complex had a labeling efficiency of approx. 100%, showed higher blood levels for longer times than conventional  $^{99m}\text{Tc}$ -albumin complexes, and was quite stable.

IT 6610-29-3

EL: ECT (Reactant); FACT (Reactant or reagent)  
(reaction of, with Et diethoxyacetate)

RN 6610-29-3 HCAPLUS

CH Hydrazinecarbothioamide, N-methyl- (PCI) (CA INDEX NAME)

S

MeNH C NH NH

L9 ANSWER 28 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1979:617698 HCAPLUS

DOCUMENT NUMBER: 91:212698

TITLE: Aqueous dispersions of copolymers with carbonyl groups and containing hydrazine derivatives

INVENTOR(S): Ley, Gregor; Penzel, Erich; Rebafka, Walter; Bott, Kaspar

PATENT ASSIGNEE(S): BASF A.-G., Fed. Rep. Ger.

SOURCE: Eur. Pat. Appl., 21 pp.

CODEN: EFXNDW

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

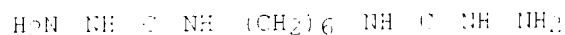
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 3510	A1	19790822	EP 1979-100168	19790119
EP 3510	B1	19810401		
R: FE, CH, DE, FR, GB, IT, NL, SE				
DE 4350070	A	19810.10	DE 1979-3465	19790116
CA 1181786	A1	19830809	CA 1979-110224	19790124
DE 7403217	A	19790727	DE 1979-317	19790125
DE 111896	B	19880111		
DE 151896	C	19880613		
NO 7901.55	A	19790727	NO 1979-255	19790125
NO 155691	B	19870303		
NO 191698	C	19870913		
ES 477135	A1	19791.01	ES 1979-477135	19790125
AT 7903557	A	19801015	AT 1979-557	19790125
AT 361536	B	19810525		
JP 54110248	A2	19790829	JP 1979-7291	19790125
JP 61066861	B4	19860391		
PRIORITY APPLN. INFO.:			DE 1978-2803253	19780126

AB Aq. coating dispersions of reaction products of polycarboxylic acid hydrazides, bis(semicarbazides), or  $\text{CO}(\text{NHNEt}_2)_2$  with aldehyde or ketone carbonyl group-contg. vinyl polymers are stabilized against hydrolysis during storage by addn. of 0.002-0.02 mol Cu, Fe, Mn, V, Sn, Cr, and/or) Ni per mol hydrazine deriv.; the metal salts are also **crosslinking** catalysts. Thus, 200 parts 10.5% aq. 25:50:25 succinic dihydrazide-glutaric dihydrazide-adipic dihydrazide dispersion and 0.06 part  $\text{CuSO}_4$  were added to a copolymer dispersion, prepd. from Me acrylate 3/5, Bu acrylate 90, acrylic acid 10, and acrolein 25 parts, to give a storage-stable dispersion. A room temp.-dried coating film swelled in DMF picking up 110-210% of its wt. in 1 day, but did not dissolve.

IT **51440-70-1D**, reaction products with carbonyl group-contg. polymers  
 FI: TEM (Technical or engineered material use); USES (Uses, coatings, stabilization of, with transition metal salts)

RN 51440-70-1 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,6-hexanediylbis- (9CI) (CA INDEX NAME)



L9 ANSWER 29 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1978:598892 HCAPLUS

DOCUMENT NUMBER: 88:193892

TITLE: Self-**crosslinkable** polyurethanes

INVENTOR(S): Winkelmann, Hans Dieter; Wolf, Karl Heinz; Oertel, Harald; Weimann, Norbert

PATENT ASSIGNEE(S): Bayer A.-G., Fed. Rep. Ger.

SOURCE: Ger. Offen., 53 pp.  
 CODEN: GWEMBK

DOCUMENT TYPE: Patent

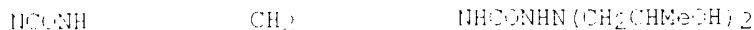
LANGUAGE: German

FAMILY APP. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 2707659	A1	19780824	DE 1977-2707659	19770223
US 4153775	A	19790508	US 1978-879504	19780111
JP 53105599	A2	19780913	JP 1978-18646	19780122
GB 1597989	A	19810916	GB 1978-7034	19780122
NL 7802036	A	19780825	NL 1978-2036	19780225
PRIORITY APPLN. INFO.:			DE 1977-2707659	19770223

GI



O

I

AB Urethane I [68125-44-0] and similar diols contg. caprolactam (II) [195-60-2]-blocked isocyanate groups were prepd. for use in the manuf. of self-**crosslinking** polyurethane elastomers. Thus, an adduct of 2

mol II and 1 mol bis(4-isocyanatophenyl)methane (III) [101-68-8] was treated with H.NN(CH<sub>2</sub>CHMeOH)<sub>2</sub> [62723-38-0] to prep. I. Adipic acid-1,6-hexanediol-neopentyl glycol copolymer (mol. wt. 1875) 500, MeN(CH<sub>2</sub>CHMeOH)<sub>2</sub> 10.58, I 37.2, and III 163.3 parts were used to prep. a prepolymer which was treated with ethylenediamine and diisocyanatohexane to prep. a **crosslinkable** copolymer [63125-45-1]. A film prepd. from the copolymer and heated at 130.degree. for 30 min was insol. in DMF at 80.degree..

IT 68125-51-9

PL: USES (Uses,

rubber, **crosslinked**)

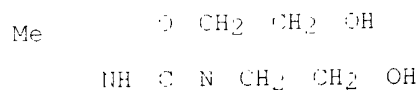
RN 68125-51-9 HCAPLUS

CN beta.-Alanine, N-(hydrazinocarbonyl)-, hydrazide, polymer with N-[3-[[[bis(2-hydroxyethyl)amino]carbonyl]amino]-4-methylphenyl]hexahydro-2-pyrimidin-1(2H)-azepine-1-carboxamide, 2,2-dimethyl-1,3-propanediol, hexanedioic acid, 1,6-hexanediol, 1,1'-methylenebis[4-isocyanatobenzene] and 1,1'-(methylimino)bis[2-propanol] (9CI) (CA INDEX NAME)

CM 1

CFN 68125-43-4

CMF C19 H28 N4 O5



NH

C O

N O

CM 2

CFN 26305-54-4

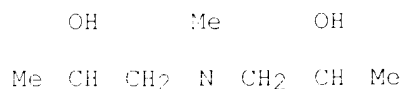
CMF C4 H11 N5 O2



CM 3

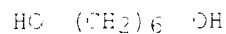
CFN 4402-30-6

CMF C7 H17 N O2



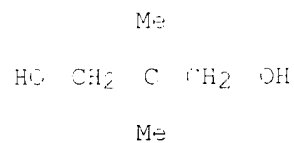
CM 4

CFN 629-11-3  
CMF CF H14 C2



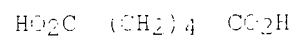
CM 5

CFN 126-30-7  
CMF CF H12 C2



CM 6

CFN 124-04-9  
CMF CF H12 C4



CM 7

CFN 101-63-8  
CMF C15 H10 N2 O2



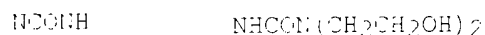
OCN

NCO

LE ANSWER 30 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1978:598891 HCAPLUS  
DOCUMENT NUMBER: 89:198891  
TITLE: Isocyanate adduct diols  
INVENTOR(S): Winkelmann, Hans Dieter; Wolf, Karl Heinz; Certel,  
Harald; Weimann, Norbert  
PATENT ASSIGNEE(S): Bayer A.-G., Fed. Rep. Ger.

SOURCE: Ger. Offen., 50 pp.  
 CODEN: GWXNBX  
 DOCUMENT TYPE: Patent  
 LANGUAGE: German  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 2707660	A1	19780824	DE 1977-2707660	19770223
DE 2707660	C2	19851214		
US 4211699	A	19800704	US 1978-879740	19780221
JP 83105428	A2	19780913	JP 1978-18647	19780222
JP 60052017	B4	19851122		
PRIORITY APPLN. INFO.:			DE 1977-2707660	19770223
GI				



AB I [68125-44-0], II [68125-48-4], and 3 similar diols were prep'd. and used for the manuf. of self-**crosslinking** polyurethane elastomers. Thus, an adduct of 2 mol caprolactam [105-60-2] and 1 mol bis(4-isocyanatophenyl)methane (III) [101-68-8] was treated with  $\text{H}_2\text{NN}(\text{CH}_2\text{CHMeOH})_2$  [62723-38-0] to prep. I. I 37.2, adipic acid-neopentyl glycol-1,6-hexanediol copolymer (mol. wt. 1375) 500,  $\text{Me}(\text{CH}_2\text{CHMeOH})_2$  16.68, and III 163.3 parts were used to prep. a prepolymer which was treated with ethylenediamine and  $\text{OCN}(\text{CH}_2)_6\text{NCO}$  to prep. a polyurethane [68125-45-1]. A film prep'd. from the polyurethane and heated at 130.degree. for 30 min was insol. in DMF at 80.degree..

IT 68125-51-9

EL: USES (Uses)  
 (rubber, **crosslinked**)

RN 68125-51-9 ECAPLUS

CN .beta.-Alanine, N-(hydrazinocarbonyl)-, hydrazide, polymer with N-[2-[[[bis(2-hydroxyethyl)amino]carbonyl]amino]-4-methylphenyl]hexahydro-2-oxo-1H-azepine-1-carboxamide, 2,2-dimethyl-1,3-propanediol, hexanedioic acid, 1,6-hexanediol, 1,1'-methylenebis[4-isocyanatobenzene] and 1,1'-(methylimino)bis[2-propanol] (9CI) (CA INDEX NAME)

CM 1

CRN 68125-48-4

DMF C19 H28 N4 O5

Me            O CH<sub>2</sub> CH<sub>2</sub> CH  
              NH C N CH<sub>2</sub> CH<sub>2</sub> OH

NH

C O

N O

CM 2

CFN 26305-54-4  
CMF C4 H11 N5 O2

O O

H<sub>2</sub>N NH C NH CH<sub>2</sub> CH<sub>2</sub> C NH NH<sub>2</sub>

CM 3

CFN 4402-30-6  
CMF C7 H17 N O2

OH Me OH

Me CH CH<sub>2</sub> N CH<sub>2</sub> CH Me

CM 4

CFN 609-11-3  
CMF C6 H14 O2

HO (CH<sub>2</sub>)<sub>6</sub> CH

CM 5

CFN 126-30-7  
CMF C5 H12 O2

Me

HO CH<sub>2</sub> C CH<sub>2</sub> OH

Me

CM 6

CFN 124-04-9  
CMF C6 H10 O4

HO<sub>2</sub>C (CH<sub>2</sub>)<sub>4</sub> CO<sub>2</sub>H

CM 7

CFN 101-64-8  
CMF C15 H10 N2 O2

CH<sub>2</sub>

GCN

NCO

L9 ANSWER 31 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1976:5924-1 HCAPLUS

DOCUMENT NUMBER: 85:192481

TITLE: Bis(thiosemicarbazido)alkanes and their main optical characteristics

AUTHOR(S): Zimenkovskii, B. S.; Turkevich, N. M.

CORPORATE SOURCE: Lvov Med. Inst., Lvov, USSR

SOURCE: Farmatsevtichnii Zhurnal (Kiev) (1976), (4), 22-6

CODEN: FFEKAP; ISSN: 0367-3057

DOCUMENT TYPE: Journal

LANGUAGE: Ukrainian

GI

O C NNHCSNH(CH<sub>2</sub>)<sub>n</sub>NNHCSNHN

N(CH<sub>2</sub>)<sub>n</sub>N

S S S S I

NMe

MeN

III

AB Hydrazinoclysis of .alpha.,.omega.-bis(thiazino)alkane derivs. I (n = 2,6) afforded H<sub>2</sub>NNHCSNH(CH<sub>2</sub>)<sub>n</sub>NNHCSNHNH<sub>2</sub> (II), which reacted with 1-methylisatin to give kisthiosemicarbazones III. II had a single uv absorption max. at

238-42 nm, corresponding to p-pi. conjugation; III had uv max. at 238-46, 273-4, and 349-54 nm, corresponding to p-pi., p-pi.\*, and the hydrazone chromophore, resp.

IT 1728-65-0P 56473-15-5P

RL: NPN (Synthetic preparation); PREF (Preparation)  
(prepn. and uv of, and reaction with methylation)

RN 1728-65-0 HCAPLUS

CN Hydrazinecarbothioamide, N,N'-1,2-ethanediyldis- (9CI) (CA INDEX NAME)

S

S

H<sub>2</sub>N NH C NH CH<sub>2</sub> CH<sub>2</sub> NH C NH NH<sub>2</sub>

RN 56473-15-5 HCAPLUS

CN Hydrazinecarbothioamide, N,N'-1,6-hexanediyldis- (9CI) (CA INDEX NAME)

S

S

H<sub>2</sub>N NH C NH (CH<sub>2</sub>)<sub>6</sub> NH C NH NH<sub>2</sub>

L9 ANSWER 32 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1975:140387 HCAPLUS

DOCUMENT NUMBER: 82:140387

TITLE: Steroid haptens

INVENTOR(S): Torelli, Vesperto; Fierdot, Andre

PATENT ASSIGNEE(S): Roussel-UCLAF

SOURCE: Ger. Offen., 45 pp.

CODEN: GWEXBX

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY APP. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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DE 2429040	A1	19750109	DE 1974-2429040	19740618
DE 2429040	C2	19851031		
EP 0035949	A1	19750131	EP 1973-20114	19730618
SE 7407108	A	19741219	SE 1974-7108	19740629
SE 402461	C	19761012		
SE 810487	A1	19741217	SE 1974-145533	19740617
NL 7408041	A	19741219	NL 1974-8041	19740617
DK 2403332	A	19750210	DK 1974-3122	19740617
US 3912191	A	19751115	US 1974-479889	19740617
CA 7408041	A	19750109	CA 1974-3641	19740617
ES 417318	A1	19760310	ES 1974-477318	19740617
BE 7404019	A0	19750111	BE 1974-4079	19740618
JP 50056451	A2	19750406	JP 1974-67309	19740618
JP 54051780	B4	19830608		
AU 740114	A1	19751219	AU 1974-70148	19740618
GE 1475356	A	19770629	GE 1974-07131	19740618
GB 1475357	A	19770629	GB 1974-3643	19740618
AT 7405045	A	19770315	AT 1974-5045	19740618
ES 448013	A1	19770701	ES 1976-448013	19760517
ES 448012	A1	19770701	ES 1976-448012	19760517



AT 351639	B	19790410	AT 1976-9863	19761210
AT 3503-63	A	14790115		
JP 5803-547	A2	14830539	JP 1982-184370	19821019
JP 5803-540	B4	14840811		
JP 5803-540	A	14830539	JP 1982-190371	19821119
JP 5803-540	B4	14830539		
JP 5803-540	A	14830539	JP 1984-75024	19840415
JP 5803-540	B4	14830539		

PRIORITY APPL. INFO.:

FR 1973-21114	19730613
AT 1974-9045	19741119

G1 For diagram(s), see printed CA Issue.

AB Estratrienols I [R = H, R1 = (CH3)3CO2H, (CH3)3CO2H; RR1 = NOCH2CO2H, NHCH2CO2H, NOCH2CO2H; R2 = H, (CH3)3CO2H; R3 = H, HO; R4 = H; R5 = HO, R4R5 = O] (10 compds.) were prepd. I [R = R1 = R3 = R4 = H; R2 = (CH3)3CO2H, R5 = HO] (II) and I [R = R1 = R3 = H, R1 = (CH3)3CO2H, R4R5 = O] formed **conjugates** with bovine serum albumins. Thus, secosteronol III was successively epoxidized, sapond., treated with CH3COCH2MgBr, hydrolyzed, cyclized, aromatized, sapond., benzylated, oxidized, treated with (EtO)3POCH2CO2Me, and hydrogenated to give I [R = R1 = R3 = R4 = H, R2 = (CH3)3CO2H, R5 = HO]. 6-Dehydro-19-nortestosterone acetate was successively treated with the tetrahydropyranyl ether of CH3COCH2OH, sapond., oxidized, and dehydrogenated to give I [R = H, R1 = (CH3)3CO2H, R2 = R3 = H, R4R5 = O].

IT 3242-64-6

FL: RCT (Reactant); RACT (Reactant or reagent)  
(reaction of, with hydroxyestratrienone)

RI 3242-64-6 HCAPLUS

CH Glycine, N-(hydrazinocarbonyl)-, monopotassium salt (9CI) (CA INDEX NAME)

Q

H,N NH C NH CH2 CO2H

● K

L6 ANSWER 53 OF 33 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1974:404780 HCAPLUS

DOCUMENT NUMBER: 51:4750

TITLE: Polyurethane coatings

INVENTOR(S): Zorn, Bruno; Noll, Klaus; Gertel, Harald; Traeubel, Harro

PATENT ASSIGNEE(S): Bayer A.-G.

SOURCE: Ger. Offen., 35 pp.

CODEN: GWXMBX

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY APP. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 2211756	A1	19731115	DE 1972-2211756	19730504
DE 2211756	B2	19791016		
DE 2211756	C3	19800620		
CA 1003712	A1	19791115	CA 1973-169961	19730425

JP 49032150	A2	19740902	CP 1973-47619	19720501
JP 48001373	B4	19830115		
IT 68-154	A	19750419	IT 1973-49340	19730502
BE 90031	A1	19731109	EE 1973-10686	19730503
NL 6806140	A	19731104	NL 1973-6180	19730503
NL 177-16	B	19350401		
NL 177-16	C	19350401		
ES 114-86	A1	19760101	ES 1973-41435	19730503
FR 1181773	A1	19731214	FR 1973-10162	19730504
GB 134-013	A	19730129	GB 1973-01174	19730504
			DE 1972-2201786	19720504

PRIORITY APPLN. INFO.:

AB Solvent-stable polyurethanes, useful as light- and abrasion-resistant coatings for textiles, leather, and leather substitutes, are prepd. by mixing solns. of aliph. or cycloaliph. diisocyanate-contg. urethane prepolymers (essentially free of NCO or NH<sub>2</sub> groups) in hydrocarbon-aliph. secondary alc. solvents with aliph. polyisocyanates, NCO functionality >2. Thus, heating adipic acid-1,4-butanediol copolymer (OH no. 31, mol. wt. 2100) 1890, OH-terminated dimethylsiloxane (OH no. 198, mol. wt. 600) 84, 1-isocyanato-3-(isocyanatomethyl)-3,5,5-trimethylcyclohexane 710, and xylene 4600 parts 2 hr at 80-100 deg. (NCO content 2.1%) and addn. of sufficient 174:4800 1-amino-3-(aminomethyl)-3,5,5-trimethylcyclohexane-MeCOOH soln. to give 25 deg. viscosity (sim. 150 P gives a soln. of clear, solid, EtOH-sol. adipic acid-1-amino-3-(aminomethyl)-3,5,5-trimethylcyclohexane-1,4-butanediol-1-isocyanato-3-(isocyanatomethyl)-3,5,5-trimethylcyclohexane copolymer [5129a-82-4]. A 12 m coating on textiles prepd. from this soln. with addn. of 30% (based on solids) com. hexamethylene diisocyanate [821-06-0]-based biuret-triisocyanate (I) cured 1 week at room temp. has very good alc. rub-fastness, compared with unsatisfactory-moderate in the presence of 0-20% I.

IT 52004-60-1

EL: TEM (Technical or engineered material use); USES (Uses)  
(coatings, for leather and textiles)

RN 52004-60-1 HCAEUS

CN .beta.-Alanine, N-(hydrazinocarbonyl)-, hydrazide, polymer with hexanedioic acid, 1,6-hexanediol and 5-isocyanato-1-(isocyanatomethyl)-1,3,5-trimethylcyclohexane (ICI) (CA INDEX NAME)

CM 1

CFN 26305-54-4  
CMF C4 H11 N5 O3

O O

H2N NH C NH CH2 CH2 C NH NH2

CM 2

CFN 4036-71-9  
CMF C12 H13 N2 O2

OCN            Me  
              CH<sub>2</sub> NCO

Me    Me

CM    2  
CPN   619-11-8  
CMF   06 H14 02

HO (CH<sub>2</sub>)<sub>6</sub> OH

CM    4  
CPN   124-04-9  
CMF   06 H10 04

HO<sub>2</sub>C (CH<sub>2</sub>)<sub>4</sub> CO<sub>2</sub>H

LA ANSWER 34 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1974:27645 HCAPLUS  
DOCUMENT NUMBER: 36:27645  
TITLE: 4,4-Alkylenebissemicarbazides and their derivatives  
INVENTOR(S): Sheppard, Chester S.; MacLeay, Ronald E.  
PATENT ASSIGNEE(S): Pennwalt Corp.  
SOURCE: U.S., 10 pp. Division of U.S. 3,585,200 (CA  
75;77759k).  
CODEN: USXXAM  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 3  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 3755238	A	19730828	US 1970-59307	19700622
US 3595200	A	19710615	US 1966-556263	19660609
PRIORITY APPLN. INFO.:			US 1966-556263	19660609

AB Substituted oxadiazolinones were treated with diamines to give alkylene bis(semicarbazides) which were used as monomers, blowing agents, and polymn. initiators. A mixt. contg. 50 g 2-phenyl-.DELTA.2-1,3,4-oxadiazolin-5-one [1109-02-6], 9.0 g ethylenediamine [107-15-3], and 250 ml H<sub>2</sub>O was refluxed 21.5 hr to give 76.5% 4,4'-ethylenebis(1-benzoylsemicarbazide) [32304-03-3], m. 263-64 deg. 4,4',4,4'-Diethylenebis-(1-benzoylsemicarbazide) [32251-24-4] was prepd. by refluxing a mixt. of 2-phenyl-.DELTA.2-1,3,4-oxadiazolin-5-one, piperazine [110-15-0] and H<sub>2</sub>O. 4,4'-Ethylenebis(semicarbazide hydrochloride) [33618-20-1] was polymd. with fumaroyl chloride [627-63-4] in 64%

yield to give a copolymer, m. .leg.300. Styrene [100-42-5] was polymd. in the presence of N,N'-ethylenebis(2-cyano-2-propylazoformamide) (I) [32251-26-9] and the rate of polymn. at 5% and 10% conversion was 6.47 .tim. 10-3 and 6.27 .tim. 10-3 moles/l.-min resp., compared to 2.81 .tim. 10-3 moles/l.-min at both conversions in the absence of I.

IT 32239-91-1P 32251-26-6P 33618-20-1P

33636-52-1P 34777-39-4P

RL: PKR (Preparation)

(prepn. of)

RN 32239-91-1 HCAPLUS

CN 2-Butenediyl dichloride, (E)-, polymer with 2,2'-(1,2-ethanediyl)bis[hydrazinecarboxamide] (9CI) (CA INDEX NAME)

CM 1

CFN 32251-26-6

CMF C4 H12 N6 O2

O

O

H2N NH C NH CH2 CH2 NH C NH NH2

CM 2

CFN 327-63-4

CMF C4 H2 Cl2 O2

Double bond geometry is shown.

O

E

Cl

Cl

O

RN 32251-26-6 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,2-ethanediylbis- (9CI) (CA INDEX NAME)

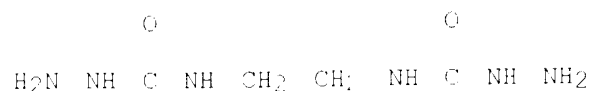
O

O

H2N NH C NH CH2 CH2 NH C NH NH2

RN 33618-20-1 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,2-ethanediylbis-, dihydrochloride (9CI) (CA INDEX NAME)

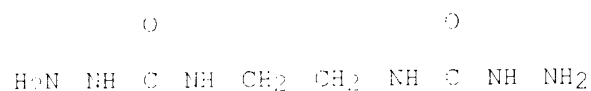


● HCl

RN 33636-52-1 HCAPLUS  
CN Carbonochloridic acid, 1,4-butanediyl ester, polymer with  
N,N'-1,2-ethanediylbis[hydrazinecarboxamide] (9CI) (CA INDEX NAME)

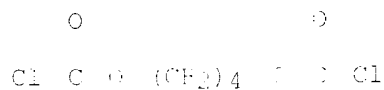
CM 1

CFN 33251-26-6  
CMF C4 H12 N6 O2



CM 2

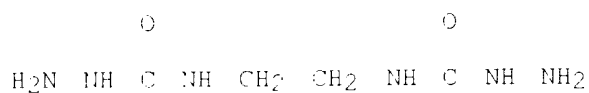
CFN 1157-16-6  
CMF C6 H8 Cl2 O4



RN 34777-39-4 HCAPLUS  
CN Hydrazinecarboxamide, N,N'-1,2-ethanediylbis-, polymer with  
2,2,4,4-tetramethyl-1,3-cyclobutanedione (9CI) (CA INDEX NAME)

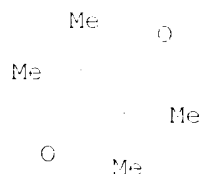
CM 1

CFN 33251-26-6  
CMF C4 H12 N6 O2



CM 2

CFN 933-52-8  
CMF C8 H12 O2



L9 ANSWER 25 OF 32 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1974:4063 HCAPLUS  
DOCUMENT NUMBER: 20:4063  
TITLE: 4,4'-Alkylenebis(semicarbazide) and derivatives  
INVENTOR(S): Sheppard, Chester G.; Macleay, Ronald E.  
PATENT ASSIGNEE(S): Pennwalt Corp.  
SOURCE: U.S., 8 pp. Division of U.S. 3,545,290 (CA 75:77759k).  
CODEN: USXXAM  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 3  
PATENT INFORMATION:

PATENT NO.	FIND	DATE	APPLICATION NO.	DATE
US 3755448	A	19710608	US 1970-50609	19700620
US 35:5200	A	19710618	US 1966-556263	19660609
PRIORITY APPLN. INFO.:			US 1966-556263	19660609

AB 4,4'-Ethylenebis(semicarbazide) (I) [32251-26-6],  
4,4'-ethylenebis(1-benzoylsemicarbazide) (II) [32304-03-3],  
4,4'-ethylenebis(semicarbazide) dihydrochloride (III) [33618-20-1]  
],  $\text{BzNHNHCOONH(CH}_2\text{)}_{12}\text{NHNHCOONHBz}$ , and 8 derivs. of I, such as  
( $\text{BzNHNHCOONHCH}_2$ )<sub>2</sub>, ( $\text{Me}_2\text{C:NHNHCOONHCH}_2$ )<sub>2</sub>, and ( $\text{NCCMe}_2\text{NHNHCOONHCH}_2$ )<sub>2</sub>, are prepd.  
Also prep'd. are the IV with R = BzNHNH, H<sub>2</sub>NNH (dihydrochloride), BzN:N,  
iso-PrOCONHNH, iso-PrOCON:N, and H<sub>2</sub>NCON:N. These compds. are used as  
monomers, polymn. catalysts, and blowing agents. Thus,  
2-phenyl-1-DELTA.,1,3,4-oxadiazolin-5-one [1199-01-6] 50, ethylenediamine  
[107-15-3] 9, and water 150 g were refluxed for 11.5 hr to prep. 76.5% II  
and a minor amt. of 4-(.beta.-aminoethyl)-1-benzoylsemicarbazide.  
Refluxing of II (3 g) in 100 ml 10% HCl for 3 3/4 hr gave 1 g III which  
was dissolved in 10 ml water and treated with 0.64 g 50% aq. NaOH to prep.  
I. A polymer of I and fumaroyl chloride (IV) [627-63-4] was prepd. by  
adding 1.53 g IV in 15 ml toluene to a soln. of III 2.43, 50% NaOH 1.6,  
NaCl 1, and Na<sub>2</sub>CO<sub>3</sub> 1.59 g in 15 ml water. This polymer was heated at  
230-50.deg., 10 mm for 2 hr to prep. a polyoxadiazole m. >300.deg..  
Refluxing of III (0.8 g) and Na acetate (0.39 g) in 13 ml water with  
tetraethyl-1,3-cyclobutanedione (V) [333-52-8] gave a I-V polymer which  
did not melt or discolor to 305.deg..

IT 32239-91-1P 33618-20-1P 33636-52-1P  
34777-39-4P

HL: PREP (Preparation)  
(prepn. of)

RN 32239-91-1 HCAPLUS

CN 2-Butyrexidoyl dichloride, (E)-, polymer with 2,2'-[1,2-ethanediylo]bis(hydrazinocarboxamide) (PCI) (CA INDEX NAME)

CM 1

GRN 32251-26-6  
CMF 34 H11 N6 02

O O  
H<sub>2</sub>N NH C NH CH<sub>2</sub> CH<sub>2</sub> NH C NH NH<sub>2</sub>

CM 2

CFN 627-63-4

CMF C4 H2 C12 O2

Double bond geometry as shown.

O  
E C1  
C1

O

RN 32613-10-1 HCAPLUS  
CN Hydrazinecarboxamide, N,N'-1,2-ethanediylbis-, dihydrochloride (9CI) (CA INDEX NAME)

O O  
H<sub>2</sub>N NH C NH CH<sub>2</sub> CH<sub>2</sub> NH C NH NH<sub>2</sub>

●2 HCl

RN 33636-50-1 HCAPLUS  
CN Carbonylchloridic acid, 1,4-butanediyl ester, polymer with  
H,N'-1,2-ethanediylbis[hydrazinecarboxamide] (9CI) (CA INDEX NAME)

CM 1

CFN 32251-16-6

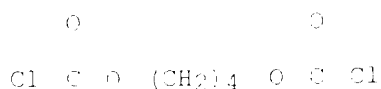
CMF C4 H12 N6 O2

O O  
H<sub>2</sub>N NH C NH CH<sub>2</sub> CH<sub>2</sub> NH C NH NH<sub>2</sub>

CM 1

CFN 1157-16-6

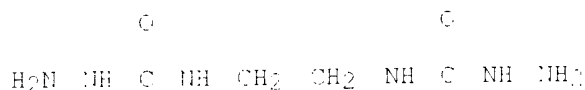
CMF C6 H8 C11 O4



RN 34777-39-4 HCAPLUS  
 CN Hydralinecarboxamide, N,N'-1,2-ethanediylbis-, polymer with  
 2,2,4,4-tetramethyl-1,3-cyclobutanedione (9CI) (CA INDEX NAME)

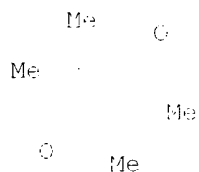
CM 1

CFN 33251-26-6  
 CMF C4 H12 N6 O2



CM 1

CFN 333-53-4  
 CMF C5 H12 N2



L9 ANSWER 36 OF 38 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1971:477759 HCAPLUS  
 DOCUMENT NUMBER: 75:77759  
 TITLE: Alkylenebis(benzoylsemicarbazides)  
 INVENTOR(S): Sheppard, Chester S.; MacLeay, Ronald E.  
 PATENT ASSIGNEE(S): Pennwalt Corp.  
 SOURCE: U.S., 8 pp.  
 CODEN: USXXAM  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 3  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 3595299	A	19710618	US 1966-556263	19660609
US 3755445	A	19730825	US 1970-59808	19700622
US 3755248	A	19730825	US 1970-59807	19700622
PRIORITY APPLN. INFO.:			US 1966-556263	19660609

GI For diagram(s), see printed CA Issue.

AB Alkylenebis(benzoylsemicarbazides), useful as intermediates in the prepn.  
 of blowing agents and polymers, were prepd. by treating  
 1-substituted-DELTA.2-1,3,4-oxadiazolin-5-ones with primary and secondary



diamines at 80-115.degree.. Thus, 50 g 2-phenyl-.DELTA.2-1,3,4-oxadiazolin-3-one and 9.0 g ethylenediamine was refluxed in 250 ml of water to give 4,4'-ethylenebis(1-benzoyl-semicarbazide) (I) m. 163-4.degree.. Similarly prepd. were 4,4',4,4'-diethylenebis(1-benzoylsemicarbazide) (II) and 4,4'-dodecamethylenebis(1-benzoylsemicarbazide). I was hydrolyzed with HCl and then treated with NaOH to yield 4,4'-ethylenebis-(semicarbazide) which copolymd. interfacially with fumaroyl chloride to give poly(fumaroyliminoureylencethyleneylenedimino) (III). On heating 2 hr at 130-50.degree./20 mm III yielded a polyoxadiazole. I.HCl was treated with H2O, NaOAc and Me2CO to yield 4,4'-ethylenebis(1-isopropylidenesemicarbazide) which was treated with HCN to give ethylenebis[1-(2-cyano-2-propyl)-semicarbazide] (IV). IV was oxidized to H,N'-ethylenebis(2-cyano-2-propylazoformamide) which initiated the polymn. of styrene and was used as a blowing agent for vinyl foams.

IT 32239-91-1P 32251-26-6P 33618-20-1P  
33636-52-1P 34777-39-4P

PL: PREP (Preparation)  
(prepn. of)

RN 32239-91-1 HCAPLUS

CN 2-Butenediyl dichloride, (E)-, polymer with 2,2'-(1,2-ethanediyl)bis(hydrazinescarboxamide) (9CI) (CA INDEX NAME)

DI 1

CFN 32251-26-6

CMF C4 H12 N6 O2

O

O

H2N NH C NH CH2 CH2 NH C NH NH2

CM 2

CFN 627-63-4

CMF C4 H2 Cl2 O2

Double bond geometry as shown.

O

E

Cl

Cl

O

FN 32251-26-6 HCAPLUS

CU Hydrazinescarboxamide, N,N'-1,2-ethanediylbis- (9CI) (CA INDEX NAME)

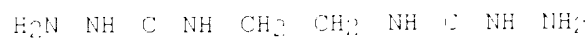
O

O

H2N NH C NH CH2 CH2 NH C NH NH2

FN 33618-20-1 HCAPLUS

CN Hydrazinecarboxamide, N,N'-1,2-ethanediylbis-, dihydrochloride (9CI) (CA INDEX NAME)



● 1 HCl

RN 33636-52-1 HCAPLUS  
CN Carbonochloridic acid, 1,4-butanediyl ester, polymer with N,N'-1,2-ethanediylbis[hydrazinecarboxamide] (9CI) (CA INDEX NAME)

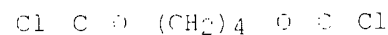
CM 1

CFN 32251-16-6  
CMF C4 H12 N6 O2



CM 1

CFN 1157-16-6  
CMF C6 H8 Cl2 O4



RN 34777-39-4 HCAPLUS  
CN Hydrazinecarboxamide, N,N'-1,2-ethanediylbis-, polymer with 2,2,4,4-tetramethyl-1,3-cyclobutanedione (9CI) (CA INDEX NAME)

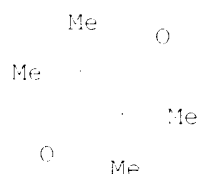
CM 1

CFN 32251-16-6  
CMF C4 H12 N6 O2



CM 1

CFN 933-52-3  
CMF C8 H12 O2



LA ANSWER 37 OF 33 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1961:70686 HCAPLUS

DOCUMENT NUMBER: 55:70686

ORIGINAL REFERENCE NO.: 55:13433a-d, 13433a-b

TITLE: Nitro olefins. II. Derivatives of .alpha.-nitroacetophenone

AUTHOR(S): Campbell, Richard D.; Schultz, Frederick J.

CORPORATE SOURCE: State Univ. of Iowa, Iowa City

SOURCE: Journal of Organic Chemistry (1960), 25, 1877-91

CODEN: JOCEAH; ISSN: 0022-0263

DOCUMENT TYPE: Journal

LANGUAGE: Unavailable

AB 25. 24. 24, 1581a. Reactions for the prepn. of series of PhC(O)CH=NO<sub>2</sub> (I) gave a series of 16 related compds. and the ultraviolet and infrared spectra were reported and discussed. I (8.25 g.) in 60 ml. dry C<sub>6</sub>H<sub>6</sub> stirred with dropwise addn. of 25 g. KOH in 40 ml. abs MeOH and the product washed twice with 1:1 MeOH-C<sub>6</sub>H<sub>6</sub> yielded 76% vacuum-dried PhC(O)CH=NO<sub>2</sub>. Similarly were prepd. PhC(O)NH<sub>4</sub>:CH=NO<sub>2</sub> and PhC(O):CH=NO<sub>2</sub> (Z = morpholinium). The prepn. of .alpha.-amino-.beta.-nitrostyrenes was accomplished by treatment of PhC(O)CH=NO<sub>2</sub> (II) with appropriate amines. PhC(tplbond)CH (10 g.) in 75 ml. cold dry Et<sub>2</sub>O treated with 15 g. liquid NOCl, the mixt. kept 10 days with gradually rising temp. and gas evolution, the pale orange liquid evapd., and the yellow oil distd. gave 16.3 g. II, m. 54-55.degree. (petr. ether), strongly lacrimatory. Et<sub>2</sub>O (75 ml.) in a heavy-wall tube cooled to -36.degree. (solid CO<sub>2</sub>-Me<sub>2</sub>SO) and bubbled through with adsorption of 15 g. NOCl, stirred with gradual addn. of 17 g. PhC(tplbond)CH and kept 2 days at -36.degree. and 7 days at 20.degree., the solvent removed in vacuo and the residue distd. gave a yellow oil, b.p. 193.5.degree., crystd. from petr. ether to yield 35.8 g. II. Freshly distd. morpholine (0.19 g.) added to 1 g. II and the Et<sub>2</sub>O-sol., H<sub>2</sub>O-insol. portion crystd. from ligroine (b. 60-70.degree.) yielded 47.5 g. .alpha.-morpholino-.beta.-nitrostyrene, m. 167-8.degree.. Under similar conditions with 2-hr. reflux of the mixt., BrNH<sub>2</sub> and II yielded 31.7 g. PhC(NHPh):CH=NO<sub>2</sub>, m. 123-4.degree., and PhCH=NH<sub>2</sub> gave 33.8 g. PhC(NHCH<sub>2</sub>Ph):CH=NO<sub>2</sub>, m. 91.degree. (CCl<sub>4</sub>). II and cyclohexylamine kept 16 hrs. yielded 65 g. PhC(NHC<sub>6</sub>H<sub>11</sub>):CH=NO<sub>2</sub>, m. 113-14.degree. (Et<sub>2</sub>O). The structures of these amine reaction products were established by acid hydrolysis to I. Several attempts were made to prep. .alpha.-acyloxy-.beta.-nitrostyrenes by acylation of I. I (4.1 g.) and 3,5-(O<sub>2</sub>N)<sub>2</sub>C<sub>6</sub>H<sub>3</sub>COCl (from 8 g. 3,5-(O<sub>2</sub>N)<sub>2</sub>C<sub>6</sub>H<sub>3</sub>COOH) refluxed 2 hrs. in 2 ml. dry C<sub>6</sub>H<sub>6</sub>N and the warm soln. filtered gave 77.4 g. PhC[3,5-(O<sub>2</sub>N)<sub>2</sub>C<sub>6</sub>H<sub>3</sub>CO]CH=NO<sub>2</sub> (III), m. 187-8.degree.. III (2.0 g.) and 25 ml. 10% NaOH warmed 3 hrs. on a steam bath and the cold soln. acidified at 0.degree. with 6M HCl, extd. with Et<sub>2</sub>O and a portion of the dried ext. chromatographed showed the presence of MeNO<sub>2</sub>. Similar acylation of I with p-O<sub>2</sub>NCH<sub>2</sub>COCl yielded 81 g. PhC(4-O<sub>2</sub>NCH<sub>2</sub>CO<sub>2</sub>)CH=NO<sub>2</sub>, m. 168-7.degree. (Me<sub>2</sub>CO, CHCl<sub>3</sub>-petr. ether). Addnl. products were obtained in a homologous series by reaction with PhCH:OMeNO<sub>2</sub> (IV) and PhCHBrOMeBrNO<sub>2</sub>. Previously were prepd. homologs PhCH:OBrNO<sub>2</sub> and PhCHBrCHBrNO<sub>2</sub>. IV (1 g.) in 15 ml. freshly distd.

morpholine kept 16 hrs. on a steam bath and the cooled soln. dild. with Et<sub>2</sub>O, washed (H<sub>2</sub>O) and evapd., the residue taken up in hot ligroine (b. 60-70.degree.) and the decolorized soln. cooled yielded 39.2% 1-phenyl-1-morpholino-2-nitropropane, m. 142-4.degree. (petr. ether). I (0.1 mole) in 175 ml. dry CH<sub>2</sub>Cl<sub>2</sub> refluxed 2 days with 0.1 mole PCl<sub>5</sub> and the residue vacuum distd. at 30.degree./12 mm., extd. with ligroine and the product recrystd. yielded 20.3% PhCHCl:O(NO<sub>2</sub>)Et, m. 90.degree.. Spectral patterns resulting from keto-enol equil., H chelation, dipole interaction, and **conjugation** effects were discussed.

IT 1728-65-0, Semicarbazide, 4,4'-ethylenedis[3-thio-  
56473-15-5, Semicarbazide, 4,4'-hexamethylenebis[3-thio-  
(prepn. of)  
BN 1728-65-0 HCAPLUS  
CN Hydrazinecarbothioamide, N,N'-1,2-ethanediylbis- (9CI) (CA INDEX NAME)

S

S

H<sub>2</sub>N NH C NH CH<sub>2</sub> CH<sub>2</sub> NH C NH NH<sub>2</sub>

BN 56473-15-5 HCAPLUS  
CN Hydrazinecarbothioamide, N,N'-1,6-hexanediylbis- (9CI) (CA INDEX NAME)

S

S

H<sub>2</sub>N NH C NH (CH<sub>2</sub>)<sub>6</sub> NH C NH NH<sub>2</sub>

LA ANSWER 38 OF 38 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1956:91004 HCAPLUS

DOCUMENT NUMBER: 56:91004

ORIGINAL REFERENCE NO.: 56:17123g-i,17124a

TITLE: The inhibition of growth of sarcoma 180 by combinations of vitamin E6 antagonists and acid hydrazides

AUTHOR(S): Brockman, F. Wallace; Thomson, J. Richard; Schabel, Frank M., Jr.; Skipper, Howard E.

CORPORATE SOURCE: Southern Research Inst., Birmingham, AL

SOURCE: Cancer Research (1956), 16, 788-95

CODEN: CNREAS; ISSN: 0008-5472

DOCUMENT TYPE: Journal

LANGUAGE: Unavailable

AB Deoxypyridoxine-HCl (I) and deoxypyridoxine phosphate (II) significantly restricted growth of sarcoma 180 in mice on a diet deficient in vitamin B6 (III), but not in mice on a complete diet. Many compds. of the acid hydrazide type also restricted growth of the sarcoma on a diet deficient in III, but none except 1,5-diaminobiuret at high dosage levels affected the tumor in mice on a complete diet. Combinations of II with acid hydrazides were more inhibitory to the tumor in mice on a complete diet than were combinations of I with acid hydrazides. The same combinations given to mice deficient in III resulted in severe restriction of tumor growth. Vitamins of the III group, i.e., pyridoxine-HCl, pyridoxamine-HCl, pyridoxal-HCl, and pyridoxal phosphate (IV), almost completely prevented the tumor-inhibiting effect of the combinations. Spectrophotometric studies demonstrated ability of the representative acid hydrazides to react with IV. The observed ability of acid hydrazides to enhance the inhibition of sarcoma 180 produced by III-deficiency and

III-antagonists is attributed to formation of an inactive  
**conjugate** between the acid hydrazides and IV.

IT 4375-11-5, Imidodicarboxylic acid, dihydrazide  
 (effect on sarcoma)  
 RN 4375-11-5 HCAPLUS  
 CN Imidodicarbonic dihydrazide (9CI) (CA INDEX NAME)

O O

H<sub>2</sub>N NH C NH C NH NH<sub>2</sub>

=&gt; d ibib abs hitstr 114 1-12

L14 ANSWER 1 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:461367 HCAPLUS

DOCUMENT NUMBER: 137:165385

TITLE: Inhibition of Cathepsin K with Lysosomotropic Macromolecular Inhibitors

AUTHOR(S): Wang, Dong; Fecar, Michal; Li, Weijie; Kopeckova, Pavla; Biedemae, Dieter; Kopecek, Jindrich

CORPORATE SOURCE: Department of Pharmaceuticals and Pharmaceutical Chemistry/OCCD and Department of Bioengineering, University of Utah, Salt Lake City, UT, 84112, USA

SOURCE: Biochemistry (2002), 41(28), 8849-8859

CODEN: BICHAU; ISSN: 0006-2960

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Cathepsin K is the major enzyme responsible for the degradn. of the protein matrix of bone and probably for the destruction of articular cartilage in rheumatoid arthritis joints. These processes occur mainly in the resorption lacuna and within the lysosomal compartment. Here, we have designed, synthesized, and evaluated new lysosomotropic (water-sol.) polymer-cathepsin K inhibitor **conjugates**. In particular, we characterized the relationship between **conjugate** structures and their activity to inhibit cathepsins K, B, L, and papain. A potent selective cathepsin K inhibitor, 1,5-bis(N-benzylloxycarbonylleucyl)carboxydranide, was modified to 1-(N-benzylloxycarbonylleucyl)-5-(phenylalanylleucyl)carboxydranide (I) to facilitate polymer **conjugation**. It was **conjugated** to the polymer chain termini of two water-sol. polymers (.alpha.-methoxy poly(ethylene glycol), abbreviated as mPEG-I; semitelechelic poly[N-(2-hydroxypropyl)methacrylamide], abbreviated as ST-PHEMA-I). The **conjugation** of inhibitor I to N-(2-hydroxypropyl)methacrylamide (HEMA) copolymer side chains was accomplished via either a Gly-Gly spacer (PHEMA-GG-I) or with no spacer between I and the copolymer backbone (PHEMA-I). Kinetic anal. revealed that free inhibitor I possessed an apparent second-order rate const. against cathepsin K ( $k_{\text{obs}}/[I] = 1.3 \times 10^6 \text{ M}^{-1} \text{ s}^{-1}$ ) similar to that of unmodified 1,5-bis(Cbz-Leu)carboxydranide, while I **conjugated** to the chain termini of mPEG and ST-PHEMA-COOH had slightly lower values (about  $3 \times 10^5 \text{ M}^{-1} \text{ s}^{-1}$ ). The  $k_{\text{obs}}/[I]$  values for I attached to the side chains of HEMA copolymers (PHEMA-GG-I and PHEMA-I) were about  $3 \times 10^4 \text{ M}^{-1} \text{ s}^{-1}$ . When tested against cathepsin L, inhibitor I and all its polymer **conjugates** produced  $k_{\text{obs}}/[I]$  values 1-2 orders of magnitude less than those detd. for cathepsin K, while for cathepsin B and papain, the values were 3-4 orders of magnitude lower. The ability of mPEG-I and ST-PHEMA-I to inhibit cathepsin K activity in synovial fibroblasts was also evaluated. Both polymer-bound inhibitors were internalized by endocytosis and were ultimately trafficked to the lysosomal compartment. ST-PHEMA-I was internalized faster than mPEG-I. The inhibitory activity in the synovial fibroblast assay correlated with the rate of internalization.

IT 190142-08-6P

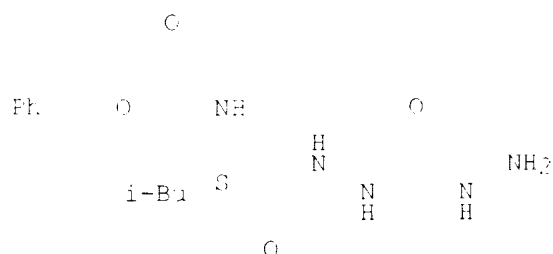
RL: RT (Reactant); SPN (Synthetic preparation); PREP (Preparation); FACT (Reactant or reagent)

(Inhibition of human lysosome cathepsin K with lysosomotropic macromol. inhibitors)

FN 190142-08-6 HCAPLUS

CN L-Leucine, N-[(phenylmethoxy)carbonyl]-, 2-(hydrazinocarbonyl)hydrazide (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 62 THERE ARE 62 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 2 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:350632 HCAPLUS

DOCUMENT NUMBER: 138:113173

TITLE: Design and synthesis of cathepsin K inhibitor-polymer **conjugates**

AUTHOR(S): Pechar, M.; Wang, D.; Kopeckova, E.; Kopecek, J.

CORPORATE SOURCE: Department of Pharmaceutics and Pharmaceutical Chemistry, University of Utah, Salt Lake City, UT, 84112, USA

SOURCE: Proceedings - 28th International Symposium on Controlled Release of Bioactive Materials and 4th Consumer & Diversified Products Conference, San Diego, CA, United States, June 23-27, 2001 (2001), Volume 2, 1319-1320. Controlled Release Society: Minneapolis, Minn.

CODEN: 69CNY8

DOCUMENT TYPE: Conference

LANGUAGE: English

AB A carbonylhydrazide based cathepsin K inhibitor was synthesized and **conjugated** with water-sol. polymers. The enzyme inhibition activities of the low mol. wt. and macromol. inhibitors were tested with papain, a model cysteine protease. The **conjugates** have the potential to facilitate delivery of the inhibitor into the bone resorption lacuna.

IT 190142-08-6DP, reaction products with polyhydroxypropylmethacrylamides

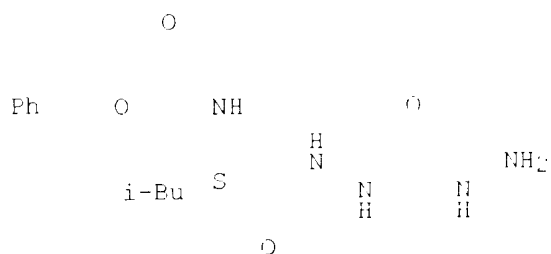
EL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(design and synthesis of cathepsin K inhibitor-polymer **conjugates**)

RN 190142-03-6 HCAPLUS

CN L-Leucine, N-[(phenylmethoxy)carbonyl]-, 2-(hydrazinocarbonyl)hydrazide (PCI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 3 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1998:684457 HCAPLUS

DOCUMENT NUMBER: 129:290447

TITLE: Preparation of branched hydrazone linkers for therapeutic drugs

INVENTOR(S): King, Dalton; Firestone, Raymond A.; Trail, Pamela

PATENT ASSIGNEE(S): USA

SOURCE: U.S., 37 pp.  
CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5314805	A	19931020	US 1996-770614	19961219
US 6511101	B1	20030128	US 1998-136351	19980819
PRIORITY APPLN. INFO.:			US 1995-9100P	P 19951122
			US 1996-770614	A3 19961219

AB Branched linkers A-Q-CONHCH[(NH)bCO-Wm-X](CH<sub>2</sub>)a(NH)bCO-(W)m-X [A is a thiol acceptor, Q is a bridging group, b and m are integers 0 or 1, W is a spacer moiety, a is an integer 2, 3, or 4, X is NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>, or NHCH[(NH)bCO-Wm-X1](CH<sub>2</sub>)a(NH)bCO-(W)m-X1, where W, a, b and are as defined, X1 is NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>, or NHCH[(NH)bCO-Wm-X2](CH<sub>2</sub>)a(NH)bCO-(W)m-X2, where W, a, b and are as defined, X2 is NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>, or NHCH[(NH)bCO-Wm-X3](CH<sub>2</sub>)a(NH)bCO-(W)m-X3, where W, a, b and are as defined, X3 is NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>, or NHCH[(NH)bCO-Wm-X4](CH<sub>2</sub>)a(NH)bCO-(W)m-X4, where W, a, b and are as defined, X4 is NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>] were prep'd. for linking a targeting ligand such as an antibody to a therapeutically active drug. Thus, the maleimidobutyrylglutamylhydrazon e of doxorubicin was prep'd. and assayed for antitumor activity.

IT 192874-02-5P

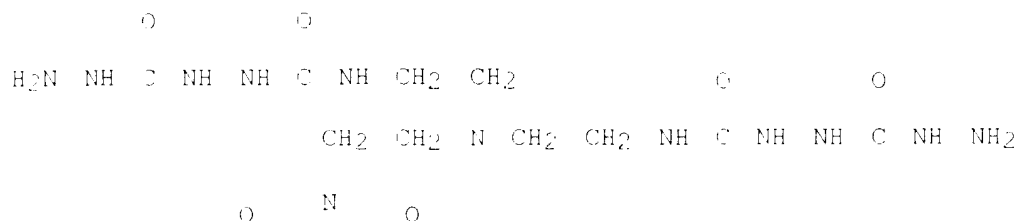
RL: SOT (Reactant); SPN (Synthetic preparation); PREP (Preparation); FACT (Reactant or reagent)

(prepn. of branched hydrazone linkers for therapeutic drugs)

RN 192874-02-5 HCAPLUS

CN 2,3,5,8,11,13,14-Heptaazapentadecanedioic acid, 3-[2-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)ethyl]-4,12-dioxo-, dihydrazide (9CI) (CA INDEX NAME)





REFERENCE COUNT: 110 THERE ARE 110 CITED REFERENCES AVAILABLE FOR  
THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE  
FORMAT

L14 ANSWER 4 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1997:503560 HCAPLUS

DOCUMENT NUMBER: 127:136079

TITLE: Preparation of branched hydrazone linkers for linking antibodies to therapeutic drugs

INVENTORS: King, Dalton; Firestone, Raymond; Trail, Pamela

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: PCT Int. Appl., 126 pp.

CODEN: PIRH2

DOCUMENT TYPE: Patent

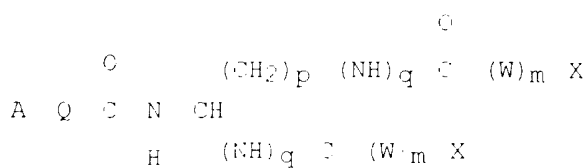
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

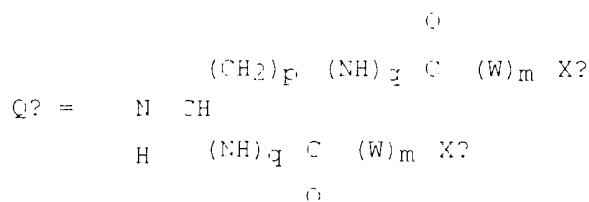
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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WO 9725243	A1	19970709	WO 1996-0320513	19961217
W: CA, JP, MX				
FW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 671490	A1	19981011	EP 1996-944522	19961217
EP 671490	B1	20030319		
F: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
JP 2000503639	T2	20000328	JP 1997-503841	19961217
AT 204635	E	20030415	AT 1996-944522	19961217
RITY APPLN. INFO.:				
			US 1995-9100P	P 19951222
			WO 1996-US20513	W 19961217

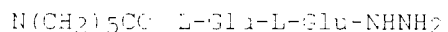
OTHER SOURCE(S):                   MARPAT 127:136079  
GI



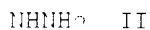
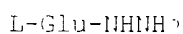
I



O



O



AB Branched hydrazone linkers I [A = thiol acceptor; Q = bridging group; q = 0, 1; W = spacer moiety; m = 0, 1; p = 2-4; X = NHNH<sub>2</sub>, moiety Q1; W, p, q, m as defined above, X1 = NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>, moiety Q2; W, p, q, m as defined above, X2 = NHNH<sub>2</sub>, moiety Q3; W, p, q, m as defined above, X3 = NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>, moiety Q4; W, p, q, m as defined above, X4 = NHNH<sub>2</sub>, NHNHCONHNH<sub>2</sub>] are claimed as agents for linking a targeting ligand such as an antibody to a therapeutically active drug. The point of branching is at a polyvalent atom and the no. of drugs increases by a factor of two for each generation of branching. A preferred drug is doxorubicin. Thus, maleimide-glutamic acid-derived hydrazone linker II was prepd. by std. coupling and deprotection methods. Condensation of II with 4 equiv of doxorubicin gave the corresponding tetrakis(hydrazone), which was then **conjugated** to monoclonal antibodies and **immunoconjugates** via the maleimide thiol acceptor. The in vivo antitumor potency and specificity of branched chain **conjugates** II and related mols. were detd.

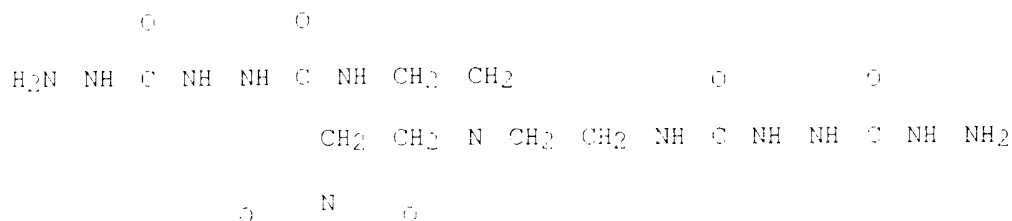
IT 192874-02-5P

PL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

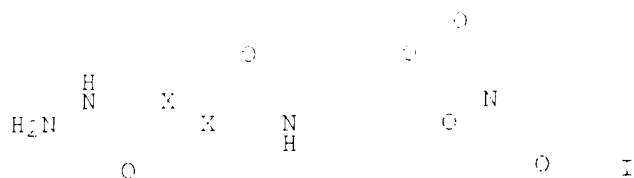
(prepn. of branched hydrazone linkers for linking antibodies to therapeutic drugs)

RN 192874-02-5 HCAPLUS

CN 2,3,5,8,11,13,14-Heptaazapentadecanedioic acid, 8-[2-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)ethyl]-4,12-dioxo-, dihydrazide (9CI) (CA INDEX NAME)

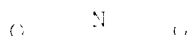
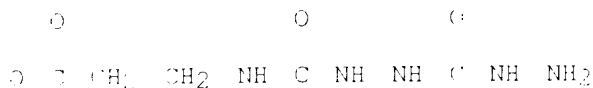


GI



CM 1

CPN 181639-49-2  
CMF 09 H14 N6 O6



CM 2

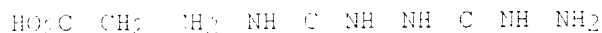
CPN 76-05-1  
CMF 02 H F3 O2

F

F C O<sub>2</sub>H

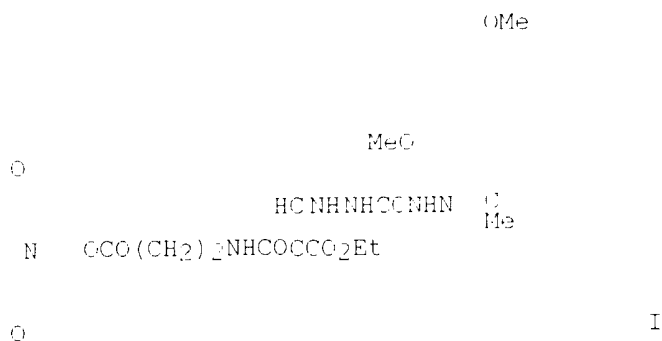
F

EN 181639-56-2 HCAPLUS  
CN .beta.-Alanine, N-[[2-(hydrazinocarbonyl)hydrazino]carbonyl]- (9CI) (CA  
INDEX NAME)



L14 ANSWER 6 OF 12 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1995:257887 HCAPLUS  
DOCUMENT NUMBER: 122:38822  
TITLE: Antibody-drug **conjugates**  
INVENTOR(S): Barton, Fussell Lavern; Briggs, Stephen Lyle  
PATENT ASSIGNEE(S): Lilly, Eli, and Co., USA  
SOURCE: Eur. Pat. Appl., 23 pp.  
CODEN: EFXXDW  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY APP. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 040044	A1	19941108	EP 1994-302952	19940425
F: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, NL, PT, SE				
JP 07 00895	A2	19950106	JP 1994-82952	19940401
CA 2121990	AA	19941029	CA 1994-2121990	19940422
PRIORITY APPLN. INFO.:			US 1993-54704	19930428
OTHER SOURCE(S):		MARPAT 122:38822		
GI				



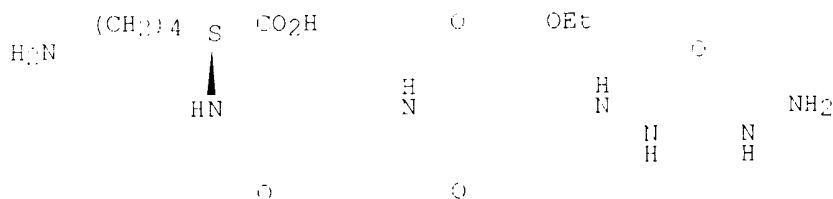
AB Malonate derivs. useful as linkers for prepn. of **immunoconjugates** comprising drugs and antibodies are provided. I was prepd. from Et malonate and .beta.-alanine benzyl ester by 6 steps and reacted with CC49 monoclonal antibody, then with doxorubicin in DMF to give an **immunoconjugate**.

IT **159795-68-3DP**, reaction products with antibody and doxorubicin  
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (prepn. of antibody-drug **conjugates**)

RN 159795-68-3 HCAPLUS

CN L-Lysine, N2-[N-[2-(ethoxycarbonyl)-3-[2-(hydrazinocarbonyl)hydrazino]-1-oxo-2-propenyl]-.beta.-alanyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.  
 Double bond geometry unknown.



L14 ANSWER 7 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1994:573214 HCAPLUS

DOCUMENT NUMBER: 121:173214

TITLE: Effect of derivatization of ribophosphate backbone and terminal ribophosphate groups in oligoribonucleotides on their stability and interaction with eukaryotic cells

AUTHOR(S): Bourtine, A. S.; Venyaminova, A. G.; Repkova, M. N.; Sergueyeva, Z. A.; Pyshnyi, D. V.

CORPORATE SOURCE: Sib. Div., Inst. Biorg. Chem., Novosibirsk, 630090, Russia

SOURCE: Biochimie (1994), 76(1), 23-32  
 CODEN: BICMBE; ISSN: 0300-9084

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Various derivs. of oligoribonucleotides were synthesized by the H-phosphonate method. Different modifications of the ribophosphate

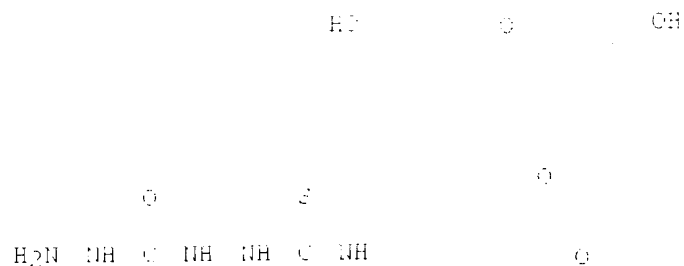
backbone were designed in order to protect the derivs. against nucleolytic enzymes present in the bio. media. These modifications include coupling of fluorescein moiety to 3'-terminal ribose, 2'-O-methylation of ribose, introduction of phosphoramidates and coupling of the last 3'-terminal nucleotide via the 3'-5'-phosphodiester bond. All modifications were tested for their effect on the stability of the derivs. against phosphodiesterase from snake venom and nucleases of the cell culture media. 2'-O-methylated oligoribonucleotides contg. either terminal 3'-5'-linkage or two 3'-terminal phosphoramidate internucleotide bonds appeared to be the most stable under the most severe conditions used. The results demonstrate a possibility to use protected oligoribonucleotide derivs. for expts. in vivo when the use of deoxy-analogs might be ineffective. The uptake of 2'-O-methylated derivs. and their 3'-cholesterol **conjugates** (coupled via a disulfide bond) by human carcinoma cells did not differ from that of the corresponding oligodeoxyribonucleotides. 95% of the bound derivs. were found in the membrane-cytosolic fraction, while only 15% were found in the nuclear fraction. The oligonucleotide moiety of 2'-O-methyloligoribonucleotide-cholesterol **conjugate** was not translocated through the cellular membrane. After cleavage of the linkage between cholesterol and oligonucleotide by dithiothreitol the major portion of the oligonucleotide moiety was released into the media. The derivs., as well as their 3'-cholesterol **conjugates**, which entered the cells, were stable and protected from action of dithiothreitol dissolved in culture media. These results demonstrate an endocytosis mechanism of penetration as obsd. in similar expts. using oligodeoxyribonucleotides.

IT 157597-83-6

EL: FCT (Reactant); FACT (Reactant or reagent)  
(reaction of, with oligoribonucleotide)

RN 157597-83-6 HCAPLUS

CN Carbonic dihydrazide, 2-[[[3',6'-dihydroxy-3-oxospiro[isobenzofuran-1(3H),9'-[9H]xanthen]-5-yl)amino]thioxomethyl]- (9CI) (CA INDEX NAME)



L14 ANSWER 8 OF 10 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1992:214841 HCAPLUS

DOCUMENT NUMBER: 116:214841

TITLE: Preparation of anthracycline **immunoconjugates**  
as neoplasm inhibitors

INVENTOR(S): Kaneko, Takusni; Willner, David; Monkovic, Ivo;  
Greenfield, Robert S.; Braslawsky, Gary R.

PATENT ASSIGNEE(S): Bristol-Myers Squibb Co., USA

SOURCE: Eur. Pat. Appl., 45 pp.

CODEN: EPXNDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 457150	A2	199111131	EP 1991-107737	19910513
EP 457150	A3	19930701		
EP 457150	B1	19960714		
E: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE				
US 5137897	A	19930811	US 1990-522895	19900514
US 5137897	B1	19960130		
AU 6174438	A1	19911114	AU 1991-74438	19910403
AU 640250	B2	19940310		
FI 2100785	A	19911115	FI 1991-21007	19910510
JP 04351765	A2	19931107	JP 1991-144757	19910510
JP 2010210	B2	20000111		
ZA 9107191	A	19920116	ZA 1991-3591	19910513
AT 183141	E	19940715	AT 1991-107737	19910513
ES 1144701	T3	19991016	ES 1991-107737	19910513
CA 2041503	AA	19911115	CA 1991-2042503	19910514
CA 2042503	C	20000713		
US 5243000	A	19940410	US 1992-565162	19920405
JP 2003026404	A2	20000125	JP 1999-181583	19990511
JP 2003026404	B2	20011204		
PRIORITY APPL. INFO.:			US 1990-512096	A 19900514
			JP 1991-149717	A3 19910510
OTHER SOURCE(S):			MARPAT 110:214841	
GI				

O OH N R1

R2  
OH

R3 O OH O

Me O

R6

R4

R5

I

AB Anthracycline derivs. I (F1 = NHCONH(CH<sub>2</sub>)<sub>n</sub>SSF<sub>8</sub>, NHCONHNHCONH(CH<sub>2</sub>)<sub>n</sub>SSF<sub>8</sub>, NHCSNH(CH<sub>2</sub>)<sub>m</sub>CH:CH(CH<sub>2</sub>)<sub>n</sub>SSF<sub>8</sub>, NHCOC(CH<sub>2</sub>)<sub>n</sub>SSF<sub>8</sub>, NHArCONH(CH<sub>2</sub>)<sub>n</sub>SSF<sub>8</sub>, etc.; m, n = 1-10; R<sub>8</sub> = (substituted) 2-pyridyl, -phenyl; Ar = phenylene; R<sub>3</sub> = Me, CH<sub>2</sub>OH, CH<sub>2</sub>OCO(CH<sub>2</sub>)<sub>3</sub>Me, CH<sub>2</sub>OCOCH(OEt)<sub>2</sub>; R<sub>3</sub> = OMe, OH, H; R<sub>4</sub> = NH<sub>2</sub>, NHCOCF<sub>3</sub>, 4-morpholinyl, 3-cyano-4-morpholinyl, 1-piperidinyl, NHCH<sub>2</sub>Ph, N(CH<sub>2</sub>Ph)<sub>2</sub>, etc.; R<sub>5</sub> = OH, tetrahydropyranyloxy, H; R<sub>6</sub> = OH, H; R<sub>6</sub> ineq. OH when R<sub>5</sub> = OH or tetrahydropyranyloxy, related compds., and their **conjugates** with ligands and antibodies, were prepd. Thus, 1-amino-4-[(2-pyridinyl)dithio]-2-butene-HCl (pregn. given) was treated with di(2-pyridyl) thionocarbonate and the product formed was condensed with Me<sub>2</sub>CO<sub>2</sub>CNHNH<sub>2</sub>. Deprotection of the resulting product by CF<sub>3</sub>CO<sub>2</sub>H gave N-[4-(2-pyridinyl)dithio]-2-butenyl]hydrazinecarbothioamide. This was condensed with adriamycin-HCl to give adriamycin 13-N-4-[(2-

pyridinyl)dithio]-2-butenylhydrazinecarbothioamide  
thiosemicarbazene,entdot.HCl (II). The **immunoconjugate** of II  
with thiolated monoclonal antibody 5E9 had IC50 of 3.0 .times. 101-7M  
against Burkitt's lymphoma cells.

IT 133701-19-6P

SL: SPN (Synthetic preparation); PREP (Preparation)  
(prepa. of, as intermediate for anticancer **immunoconjugates**)

RN 1-3701-19-6 HCAPLUS

CN Carbonic dihydrazide, 2-[[[2-(2-pyridinyl)dithio]ethyl]amino]carbonyl]-  
(-Cl) (CA INDEX NAME)

O O

N S S CH2 CH2 NH C NH NH C NH NH2

L14 ANSWER 9 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1991:426407 HCAPLUS

DOCUMENT NUMBER: 115:25407

TITLE: Novel trifunctional carrier molecule for the  
fluorescent labeling of haptens

AUTHOR(S): Bredehorst, Reinhard; Wemhoff, Gregory A.; Kusterbeck,  
Anne W.; Charles, Paul T.; Thompson, Richard B.;  
Ligler, Frances S.; Vogel, Carl Wilhelm

CORPORATE SOURCE: Dep. Biochem. Mol. Biol., Georgetown Univ.,  
Washington, DC, 20007, USA

SOURCE: Analytical Biochemistry (1991), 193(2), 272-9  
CODEN: ANBCA2; ISSN: 0003-2697

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The authors developed a novel trifunctional carrier mol. for the synthesis  
of hapten-fluorophore **conjugates** as reporter mols. in  
immunoassays. This carrier eliminates some of the disadvantages assocd.  
with currently used fluorophore-labeling procedures including high  
nonspecific binding. The backbone of the carrier consists of the Fl amino  
acid residues of the insulin A-chain mol. This polypeptide provides a  
single site (terminal amino group) for covalent coupling of the hapten,  
three carboxyl groups for the attachment of fluorophores, and four  
sulfhydryl groups for derivatization with hydrophilic residues to  
compensate for the hydrophobic effect of the attached fluorophores. The  
sites for fluorophore attachment are 4, 17, and 21 amino acids away from  
the hapten attachment site. This spatial sepn. minimizes quenching of the  
fluorescence signal due to interaction of the fluorophores with each other  
and with the attached hapten. 2,4-Dinitrophenol (DNP) was selected as  
model hapten, fluorescein as label, and S-sulfonate groups as hydrophilic  
residues. The properties of the DNP-insulin A-chain-fluorescein  
**conjugate** (DNP-Ins-Fl) were compared to those of a DNP deriv.  
labeled with a single fluorescein moiety via a small lysine spacer  
(DNP-Lys-Fl). The DNP-Ins-Fl **conjugate** exhibited a 3-fold lower  
nonspecific adsorption to **immobilized** non-immune Ig contributing  
to an approx. 3-fold more efficient displacement from the binding sites of  
an **immobilized** monoclonal anti-DNP antibody by the antigen  
DNP-lysine. Furthermore, at equimolar concns. the DNP-Ins-Fl generated a  
2.6-fold higher fluorescent signal than DNP-Lys-Fl. Due to these  
properties of DNP-Ins-Fl, DNP-lysine could be detected with an approx.



10-fold higher sensitivity compared to DNP-Lys-F1 as labeled antigen. The use of DNP-Ins-F1 as reporter molecule in a competitive fluoroimmunoassay allowed the quant. detn. of picomole amts. of DNP-lysine.

IT **134664-50-9**

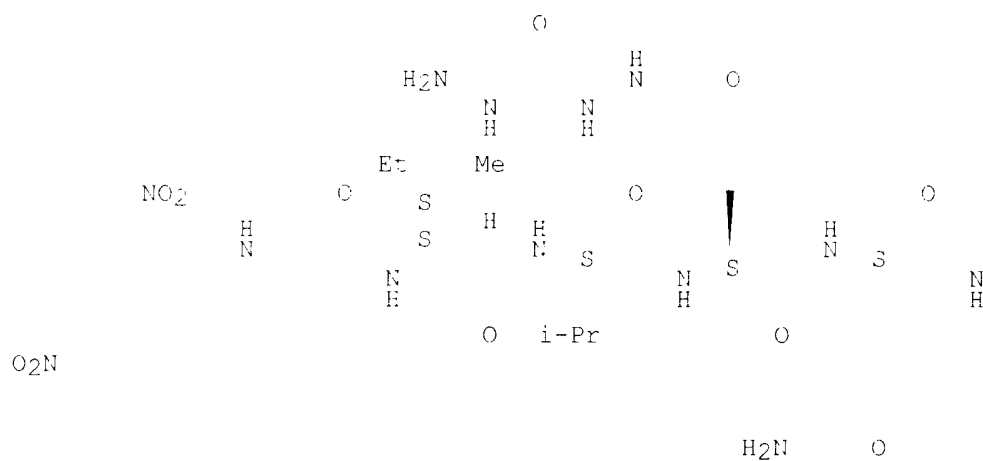
RL: RCT (Reactant); RACT (Reactant or reagent)  
(reaction of, with FITC)

RN 134664-50-9 HCAPLUS

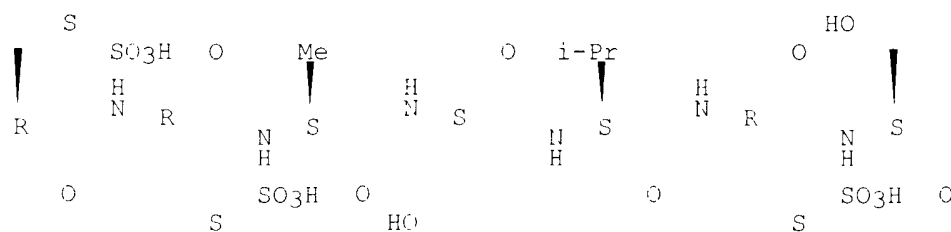
CN Insulin (cattle-A reduced), N-(2,4-dinitrophenyl)-, tris[2-(hydrazinocarbonyl)hydrazide], 6,7,11,20-tetrakis(hydrogen sulfate) (9CI)  
(CA INDEX NAME)

Absolute stereochemistry.

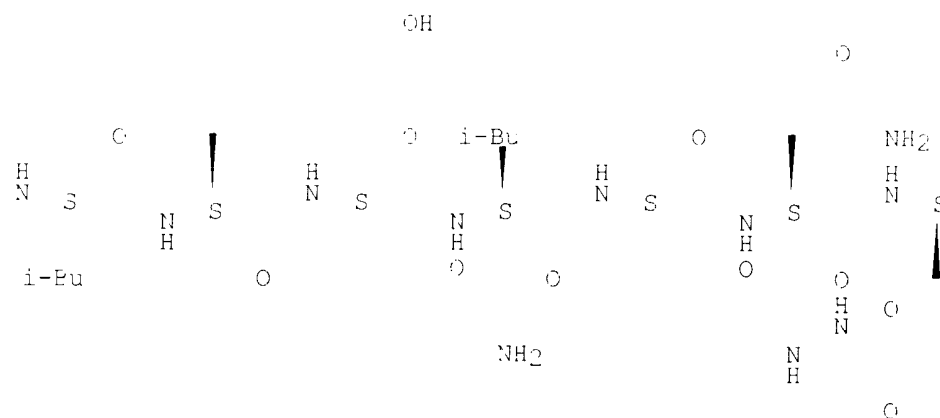
PAGE 1-A



PAGE 1-B

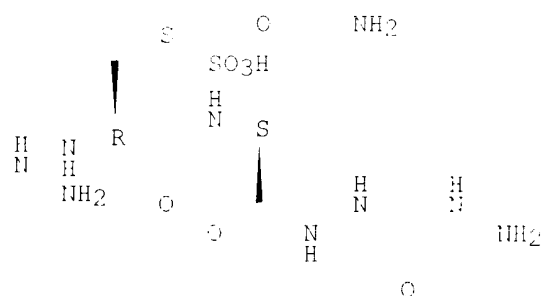


PAGE 1-C



PAGE 1-D

OH



L14 ANSWER 10 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1991:253927 HCAPLUS

DOCUMENT NUMBER: 114:253927

TITLE: New hydrazone derivatives of Adriamycin and their  
**immunoconjugates** - a correlation between acid  
stability and cytotoxicity

AUTHOR(S): Kaneko, Takushi; Willner, David; Monkovic, Ivo; Knipe,  
Jay O.; Braslawsky, Gary R.; Greenfield, Robert S.;  
Vyas, Dolatrai M.

CORPORATE SOURCE: Bristol-Myers Squibb Co., Wallingford, CT, 06492-7660,  
USA

SOURCE: Bioconjugate Chemistry (1991), 2(3), 133-41

CODEN: BOCHE5; ISSN: 1043-1802

DOCUMENT TYPE: Journal

LANGUAGE: English

AB New N-substituted hydrazine linkers were synthesized and their hydrazone derivs. of adriamycin were prepd. The adriamycin derivs. were **conjugated** with a monoclonal antibody, 5E9. The release rate of adriamycin from the hydrazones and from some of the **conjugates** was studied, and their relationship to the cytotoxicity against 5E9-pos. Daudi cells was investigated.

IT 133701-19-6P

RL: SYN (Synthetic preparation); PREP (Preparation)  
(prepn. and condensation of, with adriamycin, hydrazone from)

EN 133701-19-6 HCAPLUS

CN Carbonic dihydrazide, 2-[[[2-(2-pyridinyldithio)ethyl]amino]carbonyl]-  
(PCI) (CA INDEX NAME)

O O

N S S CH<sub>2</sub> CH<sub>2</sub> NH C NH NH C NH NH<sub>2</sub>

114 ANSWER 11 OF 12 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1983:199301 HCAPLUS

DOCUMENT NUMBER: 98:199302

TITLE: Curing of poly(glycidyl ether) resins

INVENTOR(S): Sponseller, David R.; Melky, Earl G.; Fabris, Hubert J.

PATENT ASSIGNEE(S): General Tire and Rubber Co., USA

SOURCE: U.S., 9 pp.

CODEN: USXNAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 4377680	A	19830321	US 1982-382871	19820528
PRIORITY APPLN. INFO.:			US 1982-382871	19820528

AB Cyanoalkylated hydrazides are useful as curing agents for epoxy resins, having useful pot life and showing fast cures. Thus, 1.68 g bis(cyanoethyl)carbohydrazide [85785-04-2] was mixed with 3.7 g Epon 828 [125068-38-6] to give a compn. having gel time 2.1 min at 149.degree. and room temp. pot life 6 days.

IT 85785-03-1

RL: MOA (Modifier or additive use); USES (Uses)  
(**crosslinking** agents, for epoxy resins)

EN 85785-03-1 HCAPLUS

CN Carbonic dihydrazide, 2-(2-cyanoethyl)- (PCI) (CA INDEX NAME)

O

H<sub>2</sub>N NH C NH NH CH<sub>2</sub> CH<sub>2</sub> CN

L14 ANSWER 12 OF 12 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1979:610098 HCAPLUS  
 DOCUMENT NUMBER: 01:212634  
 TITLE: Aqueous dispersions of copolymers with carbonyl groups  
 and containing hydrazine derivatives  
 INVENTOR(S): Ley, Gregor; Pensel, Erich; Fehafka, Walter; Bott,  
 Kaspar  
 PATENT ASSIGNEE(S): BASF A.-G., Fed. Rep. Ger.  
 SOURCE: Eur. Pat. Appl., 11 pp.  
 CODEN: EFXKXW  
 DOCUMENT TYPE: Patent  
 LANGUAGE: German  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 3516	A1	19790822	EP 1979-190168	19790119
EP 3516	E1	19810401		
F: EE, CH, DE, FR, GB, IT, NL, SE				
US 4150670	A	19810210	US 1979-3965	19790116
CA 1151786	A1	19830809	CA 1979-320124	19790114
DK 7300317	A	19790717	DK 1979-317	19790115
DE 111495	B	19800111		
DE 151495	C	19800113		
NO 7900255	A	19790717	NO 1979-255	19790115
NO 155695	B	19870302		
NO 155695	C	19870513		
ES 477135	A1	19791201	ES 1979-477135	19790125
AT 7900557	A	19801815	AT 1979-557	19790125
AT 361586	B	19810525		
JP 54110248	A2	19790829	JP 1979-7291	19790126
DE 61006861	B4	19860301		
PRIORITY APPLN. INFO.:		DE 1978-2803258	197-0116	

AB Aq. coating dispersions of reaction products of polycarboxylic acid  
 dihydrazides, bis(semicarbazides), or CO(NHNH2)2 with aldehyde or ketone  
 carbonyl group-contg. vinyl polymers are stabilized against hydrolysis  
 during storage by addn. of 0.0002-0.02 mol Cu, Fe, Mn, V, Zn, Cr, and(or)  
 Ni per mol hydrazine deriv.; the metal salts are also **crosslinking**  
 catalysts. Thus, 200 parts 17.5% aq. 25:50:25 succinic  
 dihydrazide-glutaric dihydrazide-adipic dihydrazide dispersion and 0.06  
 part CuSO4 were added to a copolymer dispersion, prepd. from Me acrylate  
 37%, Bu acrylate 90, acrylic acid 10, and acrolein 25 parts, to give a  
 storage-stable dispersion. A room temp.-dried coating film swelled in DMF  
 picking up 110-210% of its wt. in 1 day, but did not dissolve.

IT 1617-13-6D, reaction products with carbonyl group-contg. polymers  
 ES: TEM (Technical or engineered material use); USES (Uses)  
 (coatings, stabilization of, with transition metal salts)

RN 1617-13-6 HCAPLUS

CN 1,2-Hydrazinedicarboxylic acid, dihydrazide (9C1) (CA INDEX NAME)

0 0

H2N NH C NH NH C NH NH2

=&gt; d his

(FILE 'HOME' ENTERED AT 16:27:24 ON 06 JUN 2003)

FILE 'HCAPLUS' ENTERED AT 16:27:34 ON 06 JUN 2003

E SCHWARTZ DAVID A/AU

L1

90 S E3

L2

7 S L1 AND ?HYDRAZINE?

SELECT FN L2 2

FILE 'REGISTRY' ENTERED AT 16:28:40 ON 06 JUN 2003

L3

18 S E1-18

FILE 'HCAPLUS' ENTERED AT 16:29:17 ON 06 JUN 2003

L4

5 S L2 AND L3

FILE 'REGISTRY' ENTERED AT 16:39:27 ON 06 JUN 2003

L5

STR

L6

3 S L5

L7

609 S L5 FUL

*(Copy to see if you get L9 for structure)*

FILE 'HCAPLUS' ENTERED AT 16:41:35 ON 06 JUN 2003

L8

1059 S L7

L9

38 S L8 AND (?CROSSLINK? OR ?BIFUNCT? OR IMMOBILI? OR ?CONJUGAT?)

*25.00% from CH flow, attached*

FILE 'REGISTRY' ENTERED AT 16:43:43 ON 06 JUN 2003

L10

STR L5

L11

5 S L10

L12

95 S L10 FUL

*(Copy to see if you get L14 for structure)*

FILE 'HCAPLUS' ENTERED AT 16:46:08 ON 06 JUN 2003

L13

122 S L12

L14

12 S L13 AND (?CROSSLINK? OR ?BIFUNCT? OR IMMOBILI? OR ?CONJUGAT?)

*25.00% from CH flow, attached*

=> d que stat 19  
L5 STR

6  
G1

AK NH C NH NH2  
1 2 3 4 5

VAR G1=O/S  
NODE ATTRIBUTES:  
DEFAULT MLEVEL IS ATOM  
DEFAULT ECLEVEL IS LIMITED  
ECCOUNT IS M1-X20 C AT 1

GRAPH ATTRIBUTES:  
RING(S) ARE ISOLATED OR EMBEDDED  
NUMBER OF NODES IS 6

STEREO ATTRIBUTES: NONE  
L7 609 SEA FILE=REGISTRY SSS FUL L5  
L8 1099 SEA FILE=HCAPLUS ABB=CN L7  
L9 36 SEA FILE=HCAPLUS ABB=CN L8 AND (?CROSSLINK? OR ?BIFUNCT? OR  
IMMOBILI? OR ?CONJUGAT?)

=> d que stat l14  
L10 STR

6  
G1

AK NH NH C NH NH2  
8 7 2 3 4 5

VAR G1=O/S  
NODE ATTRIBUTES:  
DEFAULT MLEVEL IS ATOM  
DEFAULT ECLEVEL IS LIMITED  
ECCOUNT IS M1-X20 C AT 8

GRAPH ATTRIBUTES:  
RING(S) ARE ISOLATED OR EMBEDDED  
NUMBER OF NODES IS 7

STEREO ATTRIBUTES: NONE

L11 95 SEA FILE=REGISTRY SSS FUL L10  
L13 122 SEA FILE=HCAPLUS ABB=ON L12  
L14 12 SEA FILE=HCAPLUS ABB=ON L13 AND (?CROSSLINK? OR ?BIFUNCT? OR  
IMMOBILI? OR ?CONJUGAT?)